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(54) Title: CORYNEBACTERIUM GLUTAMICUM GENES ENCODING METABOLIC PATHWAY PROTEINS

(57) Abstract: Isolated nucleic acid molecules, designated MP nucleic acid molecules, which encode novel MP proteins from Corynebacterium glutamicum are described. The invention also provides antisense nucleic acid molecules, recombinant expression vectors containing MP nucleic acid molecules, and host cells into which the expression vectors have been introduced. The invention still further provides isolated MP proteins, mutated MP proteins, fusion proteins, antigenic peptides and methods for the improvement of production of a desired compound from C. glutamicum based on genetic engineering of MP genes in this organism.

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CORYNEBACTERIUM GLUTAMICUM GENES ENCODING METABOLIC PATHWAY PROTEINS

Related Applications

The present application claims priority to prior filed U.S. Provisional Patent Application Serial No. 60/141031, filed June 25, 1999, U.S. Provisional Patent 5 Application Serial No. 60/142101, filed July 2, 1999, U.S. Provisional Patent Application Serial No. 60/148613, filed August 12, 1999, and also to U.S. Provisional Patent Application Serial No. 60/187970, filed March 9, 2000. The present application also claims priority to prior filed German Patent Application No. 19930476.9, filed July 1, 1999, German Patent Application No. 19931415.2, filed July 8, 1999, German Patent 10 Application No. 19931418.7, filed July 8, 1999, German Patent Application No. 19931419.5, filed July 8, 1999, German Patent Application No. 19931420.9, filed July 8, 1999, German Patent Application No. 19931424.1, filed July 8, 1999, German Patent Application No. 19931428.4, filed July 8, 1999, German Patent Application No. 15 19931434.9, filed July 8, 1999, German Patent Application No. 19931435.7, filed July 8, 1999, German Patent Application No. 19931443.8, filed July 8, 1999, German Patent Application No. 19931453.5, filed July 8, 1999, German Patent Application No. 19931457.8, filed July 8, 1999, German Patent Application No. 19931465.9, filed July 8, 1999, German Patent Application No. 19931478.0, filed July 8, 1999, German Patent Application No. 19931510.8, filed July 8, 1999, German Patent Application No. 20 19931541.8, filed July 8, 1999, German Patent Application No. 19931573.6, filed July 8, 1999, German Patent Application No. 19931592.2, filed July 8, 1999, German Patent Application No. 19931632.3, filed July 8, 1999, German Patent Application No. 19931634.1, filed July 8, 1999, German Patent Application No. 19931636.8, filed July 25 8, 1999, German Patent Application No. 19932125.6, filed July 9, 1999, German Patent Application No. 19932126.4, filed July 9, 1999, German Patent Application No. 19932130.2, filed July 9, 1999, German Patent Application No. 19932186.8, filed July 9, 1999, German Patent Application No. 19932206.6, filed July 9, 1999, German Patent Application No. 19932227.9, filed July 9, 1999, German Patent Application No. 19932228.7, filed July 9, 1999, German Patent Application No. 19932229.5, filed July 30 9, 1999, German Patent Application No. 19932230.9, filed July 9, 1999, German Patent

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Background of the Invention

Certain products and by-products of naturally-occurring metabolic processes in cells have utility in a wide array of industries, including the food, feed, cosmetics, and pharmaceutical industries. These molecules, collectively termed 'fine chemicals', include organic acids, both proteinogenic and non-proteinogenic amino acids, nucleotides and nucleosides, lipids and fatty acids, diols, carbohydrates, aromatic compounds, vitamins and cofactors, and enzymes. Their production is most conveniently performed through large-scale culture of bacteria developed to produce and secrete large quantities of a particular desired molecule. One particularly useful organism for this purpose is *Corynebacterium glutamicum*, a gram positive, nonpathogenic bacterium. Through strain selection, a number of mutant strains have

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been developed which produce an array of desirable compounds. However, selection of strains improved for the production of a particular molecule is a time-consuming and difficult process.

5 Summary of the Invention

The invention provides novel bacterial nucleic acid molecules which have a variety of uses. These uses include the identification of microorganisms which can be used to produce fine chemicals, the modulation of fine chemical production in *C. glutamicum* or related bacteria, the typing or identification of *C. glutamicum* or related bacteria, as reference points for mapping the *C. glutamicum* genome, and as markers for transformation. These novel nucleic acid molecules encode proteins, referred to herein as metabolic pathway (MP) proteins.

C. glutamicum is a gram positive, aerobic bacterium which is commonly used in industry for the large-scale production of a variety of fine chemicals, and also for the degradation of hydrocarbons (such as in petroleum spills) and for the oxidation of terpenoids. The MP nucleic acid molecules of the invention, therefore, can be used to identify microorganisms which can be used to produce fine chemicals, e.g., by fermentation processes. Modulation of the expression of the MP nucleic acids of the invention, or modification of the sequence of the MP nucleic acid molecules of the invention, can be used to modulate the production of one or more fine chemicals from a microorganism (e.g., to improve the yield or production of one or more fine chemicals from a Corynebacterium or Brevibacterium species).

as being Corynebacterium glutamicum or a close relative thereof, or to identify the presence of C. glutamicum or a relative thereof in a mixed population of microorganisms. The invention provides the nucleic acid sequences of a number of C. glutamicum genes; by probing the extracted genomic DNA of a culture of a unique or mixed population of microorganisms under stringent conditions with a probe spanning a region of a C. glutamicum gene which is unique to this organism, one can ascertain whether this organism is present. Although Corynebacterium glutamicum itself is nonpathogenic, it is related to species pathogenic in humans, such as Corynebacterium

diphtheriae (the causative agent of diphtheria); the detection of such organisms is of significant clinical relevance.

The MP nucleic acid molecules of the invention may also serve as reference points for mapping of the C. glutamicum genome, or of genomes of related organisms. Similarly, these molecules, or variants or portions thereof, may serve as markers for genetically engineered Corynebacterium or Brevibacterium species. The MP proteins encoded by the novel nucleic acid molecules of the invention are capable of, for example, performing an enzymatic step involved in the metabolism of certain fine chemicals, including amino acids, vitamins, cofactors, nutraceuticals, 10 nucleotides, nucleosides, and trehalose. Given the availability of cloning vectors for use in Corynebacterium glutamicum, such as those disclosed in Sinskey et al., U.S. Patent No. 4,649,119, and techniques for genetic manipulation of C. glutamicum and the related Brevibacterium species (e.g., lactofermentum) (Yoshihama et al, J. Bacteriol. 162: 591-597 (1985); Katsumata et al., J. Bacteriol. 159: 306-311 (1984); and 15 Santamaria et al., J. Gen. Microbiol. 130: 2237-2246 (1984)), the nucleic acid molecules of the invention may be utilized in the genetic engineering of this organism to make it a better or more efficient producer of one or more fine chemicals.

This improved production or efficiency of production of a fine chemical may be due to a direct effect of manipulation of a gene of the invention, or it may be due to an indirect effect of such manipulation. Specifically, alterations in *C. glutamicum* metabolic pathways for amino acids, vitamins, cofactors, nucleotides, and trehalose may have a direct impact on the overall production of one or more of these desired compounds from this organism. For example, optimizing the activity of a lysine biosynthetic pathway protein or decreasing the activity of a lysine degradative pathway protein may result in an increase in the yield or efficiency of production of lysine from such an engineered organism. Alterations in the proteins involved in these metabolic pathways may also have an indirect impact on the production or efficiency of production of a desired fine chemical. For example, a reaction which is in competition for an intermediate necessary for the production of a desired molecule may be eliminated, or a pathway necessary for the production of a particular intermediate for a desired compound may be optimized. Further, modulations in the biosynthesis or degradation of, for example, an amino acid, a vitamin, or a nucleotide may increase the overall

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ability of the microorganism to rapidly grow and divide, thus increasing the number and/or production capacities of the microorganism in culture and thereby increasing the possible yield of the desired fine chemical.

The nucleic acid and protein molecules of the invention may be utilized to directly improve the production or efficiency of production of one or more desired fine chemicals from *Corynebacterium glutamicum*. Using recombinant genetic techniques well known in the art, one or more of the biosynthetic or degradative enzymes of the invention for amino acids, vitamins, cofactors, nutraceuticals, nucleotides, nucleosides, or trehalose may be manipulated such that its function is modulated. For example, a biosynthetic enzyme may be improved in efficiency, or its allosteric control region destroyed such that feedback inhibition of production of the compound is prevented. Similarly, a degradative enzyme may be deleted or modified by substitution, deletion, or addition such that its degradative activity is lessened for the desired compound without impairing the viability of the cell. In each case, the overall yield or rate of production of the desired fine chemical may be increased.

It is also possible that such alterations in the protein and nucleotide molecules of the invention may improve the production of other fine chemicals besides the amino acids, vitamins, cofactors, nutraceuticals, nucleotides, nucleosides, and trehalose through indirect mechanisms. Metabolism of any one compound is necessarily intertwined with other biosynthetic and degradative pathways within the cell, and necessary cofactors, intermediates, or substrates in one pathway are likely supplied or limited by another such pathway. Therefore, by modulating the activity of one or more of the proteins of the invention, the production or efficiency of activity of another fine chemical biosynthetic or degradative pathway may be impacted. For example, amino acids serve as the structural units of all proteins, yet may be present intracellularly in levels which are limiting for protein synthesis; therefore, by increasing the efficiency of production or the yields of one or more amino acids within the cell, proteins, such as biosynthetic or degradative proteins, may be more readily synthesized. Likewise, an alteration in a metabolic pathway enzyme such that a particular side reaction becomes more or less favored may result in the over- or under-production of one or more compounds which are utilized as intermediates or substrates for the production of a desired fine chemical.

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This invention provides novel nucleic acid molecules which encode proteins, referred to herein as metabolic pathway proteins (MP), which are capable of, for example, performing an enzymatic step involved in the metabolism of molecules important for the normal functioning of cells, such as amino acids, vitamins, cofactors, nucleotides and nucleosides, or trehalose. Nucleic acid molecules encoding an MP protein are referred to herein as MP nucleic acid molecules. In a preferred embodiment, the MP protein performs an enzymatic step related to the metabolism of one or more of the following: amino acids, vitamins, cofactors, nutraceuticals, nucleotides, nucleosides, and trehalose. Examples of such proteins include those encoded by the genes set forth in Table 1.

Accordingly, one aspect of the invention pertains to isolated nucleic acid molecules (e.g., cDNAs, DNAs, or RNAs) comprising a nucleotide sequence encoding an MP protein or biologically active portions thereof, as well as nucleic acid fragments suitable as primers or hybridization probes for the detection or amplification of MPencoding nucleic acid (e.g., DNA or mRNA). In particularly preferred embodiments, the isolated nucleic acid molecule comprises one of the nucleotide sequences set forth as the odd-numbered SEQ ID NOs in the Sequence Listing (e.g., SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:5, SEQ ID NO:7....), or the coding region or a complement thereof of one of these nucleotide sequences. In other particularly preferred embodiments, the isolated nucleic acid molecule of the invention comprises a nucleotide sequence which hybridizes to or is at least about 50%, preferably at least about 60%, more preferably at least about 70%, 80% or 90%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more homologous to a nucleotide sequence set forth as an odd-numbered SEQ ID NO in the Sequence Listing (e.g., SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:5, SEQ ID NO:7....), or a portion thereof. In other preferred embodiments, the isolated nucleic acid molecule encodes one of the amino acid sequences set forth as an evennumbered SEQ ID NO in the Sequence Listing (e.g., SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:6, SEQ ID NO:8....). The preferred MP proteins of the present invention also preferably possess at least one of the MP activities described herein.

In another embodiment, the isolated nucleic acid molecule encodes a protein or portion thereof wherein the protein or portion thereof includes an amino acid sequence which is sufficiently homologous to an amino acid sequence of the invention (e.g., a

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sequence having an even-numbered SEQ ID NO: in the Sequence Listing), *e.g.*, sufficiently homologous to an amino acid sequence of the invention such that the protein or portion thereof maintains an MP activity. Preferably, the protein or portion thereof encoded by the nucleic acid molecule maintains the ability to perform an enzymatic reaction in a amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathway. In one embodiment, the protein encoded by the nucleic acid molecule is at least about 50%, preferably at least about 60%, and more preferably at least about 70%, 80%, or 90% and most preferably at least about 95%, 96%, 97%, 98%, or 99% or more homologous to an amino acid sequence of the invention (*e.g.*, an entire amino acid sequence selected from those having an even-numbered SEQ ID NO in the Sequence Listing). In another preferred embodiment, the protein is a full length *C. glutamicum* protein which is substantially homologous to an entire amino acid sequence of the invention (encoded by an open reading frame shown in the corresponding odd-numbered SEQ ID NOs in the Sequence Listing (*e.g.*, SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:5, SEQ ID NO:7....).

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In another preferred embodiment, the isolated nucleic acid molecule is derived from *C. glutamicum* and encodes a protein (*e.g.*, an MP fusion protein) which includes a biologically active domain which is at least about 50% or more homologous to one of the amino acid sequences of the invention (*e.g.*, a sequence of one of the even-numbered SEQ ID NOs in the Sequence Listing) and is able to catalyze a reaction in a metabolic pathway for an amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose, or one or more of the activities set forth in Table 1, and which also includes heterologous nucleic acid sequences encoding a heterologous polypeptide or regulatory regions.

In another embodiment, the isolated nucleic acid molecule is at least 15 nucleotides in length and hybridizes under stringent conditions to a nucleic acid molecule comprising a nucleotide sequence of the invention (e.g., a sequence of an odd-numbered SEQ ID NO in the Sequence Listing). Preferably, the isolated nucleic acid molecule corresponds to a naturally-occurring nucleic acid molecule. More preferably, the isolated nucleic acid encodes a naturally-occurring *C. glutamicum* MP protein, or a biologically active portion thereof.

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Another aspect of the invention pertains to vectors, *e.g.*, recombinant expression vectors, containing the nucleic acid molecules of the invention, and host cells into which such vectors have been introduced. In one embodiment, such a host cell is used to produce an MP protein by culturing the host cell in a suitable medium. The MP protein can be then isolated from the medium or the host cell.

Yet another aspect of the invention pertains to a genetically altered microorganism in which an MP gene has been introduced or altered. In one embodiment, the genome of the microorganism has been altered by introduction of a nucleic acid molecule of the invention encoding wild-type or mutated MP sequence as a transgene. In another embodiment, an endogenous MP gene within the genome of the microorganism has been altered, e.g., functionally disrupted, by homologous recombination with an altered MP gene. In another embodiment, an endogenous or introduced MP gene in a microorganism has been altered by one or more point mutations, deletions, or inversions, but still encodes a functional MP protein. In still another embodiment, one or more of the regulatory regions (e.g., a promoter, repressor, or inducer) of an MP gene in a microorganism has been altered (e.g., by deletion, truncation, inversion, or point mutation) such that the expression of the MP gene is modulated. In a preferred embodiment, the microorganism belongs to the genus Corynebacterium or Brevibacterium, with Corynebacterium glutamicum being particularly preferred. In a preferred embodiment, the microorganism is also utilized for the production of a desired compound, such as an amino acid, with lysine being particularly preferred.

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In another aspect, the invention provides a method of identifying the presence or activity of *Cornyebacterium diphtheriae* in a subject. This method includes detection of one or more of the nucleic acid or amino acid sequences of the invention (*e.g.*, the sequences set forth in the Sequence Listing as SEQ ID NOs 1 through 1156) in a subject, thereby detecting the presence or activity of *Corynebacterium diphtheriae* in the subject.

Still another aspect of the invention pertains to an isolated MP protein or a portion, *e.g.*, a biologically active portion, thereof. In a preferred embodiment, the isolated MP protein or portion thereof can catalyze an enzymatic reaction involved in one or more pathways for the metabolism of an amino acid, a vitamin, a cofactor, a

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nutraceutical, a nucleotide, a nucleoside, or trehalose. In another preferred embodiment, the isolated MP protein or portion thereof is sufficiently homologous to an amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: in the Sequence Listing) such that the protein or portion thereof maintains the ability to catalyze an enzymatic reaction involved in one or more pathways for the metabolism of an amino acid, a vitamin, a cofactor, a nutraceutical, a nucleotide, a nucleoside, or trehalose.

The invention also provides an isolated preparation of an MP protein. In preferred embodiments, the MP protein comprises an amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing). In another preferred embodiment, the invention pertains to an isolated full length protein which is substantially homologous to an entire amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing) (encoded by an open reading frame set forth in a corresponding odd-numbered SEQ ID NO: of the Sequence Listing). In yet another embodiment, the protein is at least about 50%, preferably at least about 60%, and more preferably at least about 70%, 80%, or 90%, and most preferably at least about 95%, 96%, 97%, 98%, or 99% or more homologous to an entire amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing). In other embodiments, the isolated MP protein comprises an amino acid sequence which is at least about 50% or more homologous to one of the amino acid sequences of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing) and is able to catalyze an enzymatic reaction in an amino acid, vitamin, cofactor, nutraccutical, nucleotide, nucleoside, or trehalose metabolic pathway, or has one or more of the activities set forth in Table 1.

Alternatively, the isolated MP protein can comprise an amino acid sequence which is encoded by a nucleotide sequence which hybridizes, *e.g.*, hybridizes under stringent conditions, or is at least about 50%, preferably at least about 60%, more preferably at least about 70%, 80%, or 90%, and even more preferably at least about 95%, 96%, 97%, 98,%, or 99% or more homologous to a nucleotide sequence of one of the even-numbered SEQ ID NOs set forth in the Sequence Listing. It is also preferred that the preferred forms of MP proteins also have one or more of the MP bioactivities described herein.

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The MP polypeptide, or a biologically active portion thereof, can be operatively linked to a non-MP polypeptide to form a fusion protein. In preferred embodiments, this fusion protein has an activity which differs from that of the MP protein alone. In other preferred embodiments, this fusion protein, when introduced into a *C. glutamicum* pathway for the metabolism of an amino acid, vitamin, cofactor, nutraceutical, results in increased yields and/or efficiency of production of a desired fine chemical from *C. glutamicum*. In particularly preferred embodiments, integration of this fusion protein into an amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathway of a host cell modulates production of a desired compound from the cell.

In another aspect, the invention provides methods for screening molecules which modulate the activity of an MP protein, either by interacting with the protein itself or a substrate or binding partner of the MP protein, or by modulating the transcription or translation of an MP nucleic acid molecule of the invention.

Another aspect of the invention pertains to a method for producing a fine chemical. This method involves the culturing of a cell containing a vector directing the expression of an MP nucleic acid molecule of the invention, such that a fine chemical is produced. In a preferred embodiment, this method further includes the step of obtaining a cell containing such a vector, in which a cell is transfected with a vector directing the expression of an MP nucleic acid. In another preferred embodiment, this method further includes the step of recovering the fine chemical from the culture. In a particularly preferred embodiment, the cell is from the genus *Corynebacterium* or *Brevibacterium*, or is selected from those strains set forth in Table 3.

Another aspect of the invention pertains to methods for modulating production of a molecule from a microorganism. Such methods include contacting the cell with an agent which modulates MP protein activity or MP nucleic acid expression such that a cell associated activity is altered relative to this same activity in the absence of the agent. In a preferred embodiment, the cell is modulated for one or more *C. glutamicum* amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathways, such that the yields or rate of production of a desired fine chemical by this microorganism is improved. The agent which modulates MP protein activity can be an agent which stimulates MP protein activity or MP nucleic acid expression.

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Examples of agents which stimulate MP protein activity or MP nucleic acid expression include small molecules, active MP proteins, and nucleic acids encoding MP proteins that have been introduced into the cell. Examples of agents which inhibit MP activity or expression include small molecules, and antisense MP nucleic acid molecules.

Another aspect of the invention pertains to methods for modulating yields of a desired compound from a cell, involving the introduction of a wild-type or mutant MP gene into a cell, either maintained on a separate plasmid or integrated into the genome of the host cell. If integrated into the genome, such integration can be random, or it can take place by homologous recombination such that the native gene is replaced by the introduced copy, causing the production of the desired compound from the cell to be modulated. In a preferred embodiment, said yields are increased. In another preferred embodiment, said chemical is a fine chemical. In a particularly preferred embodiment, said fine chemical is an amino acid. In especially preferred embodiments, said amino acid is L-lysine.

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Detailed Description of the Invention

The present invention provides MP nucleic acid and protein molecules which are involved in the metabolism of certain fine chemicals in *Corynebacterium glutamicum*, including amino acids, vitamins, cofactors, nutraceuticals, nucleotides, nucleosides, and trehalose. The molecules of the invention may be utilized in the modulation of production of fine chemicals from microorganisms, such as *C. glutamicum*, either directly (*e.g.*, where modulation of the activity of a lysine biosynthesis protein has a direct impact on the production or efficiency of production of lysine from that organism), or may have an indirect impact which nonetheless results in an increase of yield or efficiency of production of the desired compound (*e.g.*, where modulation of the activity of a nucleotide biosynthesis protein has an impact on the production of an organic acid or a fatty acid from the bacterium, perhaps due to improved growth or an increased supply of necessary co-factors, energy compounds, or precursor molecules). Aspects of the invention are further explicated below.

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I. Fine Chemicals

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The term 'fine chemical' is art-recognized and includes molecules produced by an organism which have applications in various industries, such as, but not limited to, the pharmaceutical, agriculture, and cosmetics industries. Such compounds include organic acids, such as tartaric acid, itaconic acid, and diaminopimelic acid, both proteinogenic and non-proteinogenic amino acids, purine and pyrimidine bases, nucleosides, and nucleotides (as described e.g. in Kuninaka, A. (1996) Nucleotides and related compounds, p. 561-612, in Biotechnology vol. 6, Rehm et al., eds. VCH: Weinheim, and references contained therein), lipids, both saturated and unsaturated fatty acids (e.g., arachidonic acid), diols (e.g., propane diol, and butane diol), carbohydrates 10 (e.g., hyaluronic acid and trehalose), aromatic compounds (e.g., aromatic amines, vanillin, and indigo), vitamins and cofactors (as described in Ullmann's Encyclopedia of Industrial Chemistry, vol. A27, "Vitamins", p. 443-613 (1996) VCH: Weinheim and references therein; and Ong, A.S., Niki, E. & Packer, L. (1995) "Nutrition, Lipids, Health, and Disease" Proceedings of the UNESCO/Confederation of Scientific and 15 Technological Associations in Malaysia, and the Society for Free Radical Research -Asia, held Sept. 1-3, 1994 at Penang, Malaysia, AOCS Press, (1995)), enzymes, polyketides (Cane et al. (1998) Science 282: 63-68), and all other chemicals described in Gutcho (1983) Chemicals by Fermentation, Noyes Data Corporation, ISBN: 0818805086 and references therein. The metabolism and uses of certain of these fine 20 chemicals are further explicated below.

A. Amino Acid Metabolism and Uses

Amino acids comprise the basic structural units of all proteins, and as such are
essential for normal cellular functioning in all organisms. The term "amino acid" is artrecognized. The proteinogenic amino acids, of which there are 20 species, serve as
structural units for proteins, in which they are linked by peptide bonds, while the
nonproteinogenic amino acids (hundreds of which are known) are not normally found in
proteins (see Ulmann's Encyclopedia of Industrial Chemistry, vol. A2, p. 57-97 VCH:
Weinheim (1985)). Amino acids may be in the D- or L- optical configuration, though Lamino acids are generally the only type found in naturally-occurring proteins.
Biosynthetic and degradative pathways of each of the 20 proteinogenic amino acids

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have been well characterized in both prokaryotic and eukaryotic cells (see, for example, Stryer, L. Biochemistry, 3rd edition, pages 578-590 (1988)). The 'essential' amino acids (histidine, isoleucine, leucine, lysine, methionine, phenylalanine, threonine, tryptophan, and valine), so named because they are generally a nutritional requirement due to the complexity of their biosyntheses, are readily converted by simple biosynthetic pathways to the remaining 11 'nonessential' amino acids (alanine, arginine, asparagine, aspartate, cysteine, glutamate, glutamine, glycine, proline, serine, and tyrosine). Higher animals do retain the ability to synthesize some of these amino acids, but the essential amino acids must be supplied from the diet in order for normal protein synthesis to occur.

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Aside from their function in protein biosynthesis, these amino acids are interesting chemicals in their own right, and many have been found to have various applications in the food, feed, chemical, cosmetics, agriculture, and pharmaceutical industries. Lysine is an important amino acid in the nutrition not only of humans, but also of monogastric animals such as poultry and swine. Glutamate is most commonly used as a flavor additive (mono-sodium glutamate, MSG) and is widely used throughout the food industry, as are aspartate, phenylalanine, glycine, and cysteine. Glycine, Lmethionine and tryptophan are all utilized in the pharmaceutical industry. Glutamine, valine, leucine, isoleucine, histidine, arginine, proline, serine and alanine are of use in both the pharmaceutical and cosmetics industries. Threonine, tryptophan, and D/Lmethionine are common feed additives. (Leuchtenberger, W. (1996) Amino aids – technical production and use, p. 466-502 in Rehm et al. (eds.) Biotechnology vol. 6, chapter 14a, VCH: Weinheim). Additionally, these amino acids have been found to be useful as precursors for the synthesis of synthetic amino acids and proteins, such as Nacetylcysteine, S-carboxymethyl-L-cysteine, (S)-5-hydroxytryptophan, and others described in Ulmann's Encyclopedia of Industrial Chemistry, vol. A2, p. 57-97, VCH: Weinheim, 1985.

The biosynthesis of these natural amino acids in organisms capable of producing them, such as bacteria, has been well characterized (for review of bacterial amino acid biosynthesis and regulation thereof, see Umbarger, H.E.(1978) *Ann. Rev. Biochem.* 47: 533-606). Glutamate is synthesized by the reductive amination of α-ketoglutarate, an intermediate in the citric acid cycle. Glutamine, proline, and arginine are each subsequently produced from glutamate. The biosynthesis of serine is a three-

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step process beginning with 3-phosphoglycerate (an intermediate in glycolysis), and resulting in this amino acid after oxidation, transamination, and hydrolysis steps. Both cysteine and glycine are produced from serine; the former by the condensation of homocysteine with serine, and the latter by the transferal of the side-chain β -carbon atom to tetrahydrofolate, in a reaction catalyzed by serine transhydroxymethylase. Phenylalanine, and tyrosine are synthesized from the glycolytic and pentose phosphate pathway precursors erythrose 4-phosphate and phosphoenolpyruvate in a 9-step biosynthetic pathway that differ only at the final two steps after synthesis of prephenate. Tryptophan is also produced from these two initial molecules, but its synthesis is an 11step pathway. Tyrosine may also be synthesized from phenylalanine, in a reaction catalyzed by phenylalanine hydroxylase. Alanine, valine, and leucine are all biosynthetic products of pyruvate, the final product of glycolysis. Aspartate is formed from oxaloacetate, an intermediate of the citric acid cycle. Asparagine, methionine, threonine, and lysine are each produced by the conversion of aspartate. Isoleucine is formed from threonine. A complex 9-step pathway results in the production of histidine 15 from 5-phosphoribosyl-1-pyrophosphate, an activated sugar.

Amino acids in excess of the protein synthesis needs of the cell cannot be stored, and are instead degraded to provide intermediates for the major metabolic pathways of the cell (for review see Stryer, L. Biochemistry 3rd ed. Ch. 21 "Amino Acid Degradation and the Urea Cycle" p. 495-516 (1988)). Although the cell is able to convert unwanted amino acids into useful metabolic intermediates, amino acid production is costly in terms of energy, precursor molecules, and the enzymes necessary to synthesize them. Thus it is not surprising that amino acid biosynthesis is regulated by feedback inhibition, in which the presence of a particular amino acid serves to slow or entirely stop its own production (for overview of feedback mechanisms in amino acid biosynthetic pathways, see Stryer, L. Biochemistry, 3rd ed. Ch. 24: "Biosynthesis of Amino Acids and Heme" p. 575-600 (1988)). Thus, the output of any particular amino acid is limited by the amount of that amino acid present in the cell.

B. Vitamin, Cofactor, and Nutraceutical Metabolism and Uses 30

Vitamins, cofactors, and nutraceuticals comprise another group of molecules which the higher animals have lost the ability to synthesize and so must ingest, although

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they are readily synthesized by other organisms, such as bacteria. These molecules are either bioactive substances themselves, or are precursors of biologically active substances which may serve as electron carriers or intermediates in a variety of metabolic pathways. Aside from their nutritive value, these compounds also have significant industrial value as coloring agents, antioxidants, and catalysts or other processing aids. (For an overview of the structure, activity, and industrial applications of these compounds, see, for example, Ullman's Encyclopedia of Industrial Chemistry, "Vitamins" vol. A27, p. 443-613, VCH: Weinheim, 1996.) The term "vitamin" is artrecognized, and includes nutrients which are required by an organism for normal functioning, but which that organism cannot synthesize by itself. The group of vitamins may encompass cofactors and nutraceutical compounds. The language "cofactor" includes nonproteinaceous compounds required for a normal enzymatic activity to occur. Such compounds may be organic or inorganic; the cofactor molecules of the invention are preferably organic. The term "nutraceutical" includes dietary supplements having health benefits in plants and animals, particularly humans. Examples of such molecules are vitamins, antioxidants, and also certain lipids (e.g., polyunsaturated fatty acids).

The biosynthesis of these molecules in organisms capable of producing them, such as bacteria, has been largely characterized (Ullman's Encyclopedia of Industrial Chemistry, "Vitamins" vol. A27, p. 443-613, VCH: Weinheim, 1996; Michal, G. (1999) Biochemical Pathways: An Atlas of Biochemistry and Molecular Biology, John Wiley & Sons; Ong, A.S., Niki, E. & Packer, L. (1995) "Nutrition, Lipids, Health, and Disease" Proceedings of the UNESCO/Confederation of Scientific and Technological Associations in Malaysia, and the Society for Free Radical Research – Asia, held Sept. 1-3, 1994 at Penang, Malaysia, AOCS Press: Champaign, IL X, 374 S).

Thiamin (vitamin B₁) is produced by the chemical coupling of pyrimidine and thiazole moieties. Riboflavin (vitamin B₂) is synthesized from guanosine-5'-triphosphate (GTP) and ribose-5'-phosphate. Riboflavin, in turn, is utilized for the synthesis of flavin mononucleotide (FMN) and flavin adenine dinucleotide (FAD). The family of compounds collectively termed 'vitamin B₆' (e.g., pyridoxine, pyridoxamine, pyridoxa-5'-phosphate, and the commercially used pyridoxin hydrochloride) are all derivatives of the common structural unit, 5-hydroxy-6-methylpyridine. Pantothenate (pantothenic

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acid, (R)-(+)-N-(2,4-dihydroxy-3,3-dimethyl-1-oxobutyl)- β -alanine) can be produced either by chemical synthesis or by fermentation. The final steps in pantothenate biosynthesis consist of the ATP-driven condensation of β -alanine and pantoic acid. The enzymes responsible for the biosynthesis steps for the conversion to pantoic acid, to β -alanine and for the condensation to panthotenic acid are known. The metabolically active form of pantothenate is Coenzyme A, for which the biosynthesis proceeds in 5 enzymatic steps. Pantothenate, pyridoxal-5'-phosphate, cysteine and ATP are the precursors of Coenzyme A. These enzymes not only catalyze the formation of panthothante, but also the production of (R)-pantoic acid, (R)-pantolacton, (R)-panthenol (provitamin B₅), pantetheine (and its derivatives) and coenzyme A.

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Biotin biosynthesis from the precursor molecule pimeloyl-CoA in microorganisms has been studied in detail and several of the genes involved have been identified. Many of the corresponding proteins have been found to also be involved in Fe-cluster synthesis and are members of the nifS class of proteins. Lipoic acid is derived from octanoic acid, and serves as a coenzyme in energy metabolism, where it becomes part of the pyruvate dehydrogenase complex and the α-ketoglutarate dehydrogenase complex. The folates are a group of substances which are all derivatives of folic acid, which is turn is derived from L-glutamic acid, p-amino-benzoic acid and 6-methylpterin. The biosynthesis of folic acid and its derivatives, starting from the metabolism intermediates guanosine-5'-triphosphate (GTP), L-glutamic acid and p-amino-benzoic acid has been studied in detail in certain microorganisms.

Corrinoids (such as the cobalamines and particularly vitamin B₁₂) and porphyrines belong to a group of chemicals characterized by a tetrapyrole ring system. The biosynthesis of vitamin B₁₂ is sufficiently complex that it has not yet been completely characterized, but many of the enzymes and substrates involved are now known. Nicotinic acid (nicotinate), and nicotinamide are pyridine derivatives which are also termed 'niacin'. Niacin is the precursor of the important coenzymes NAD (nicotinamide adenine dinucleotide) and NADP (nicotinamide adenine dinucleotide phosphate) and their reduced forms.

The large-scale production of these compounds has largely relied on cell-free chemical syntheses, though some of these chemicals have also been produced by large-scale culture of microorganisms, such as riboflavin, Vitamin B₆, pantothenate, and

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biotin. Only Vitamin B_{12} is produced solely by fermentation, due to the complexity of its synthesis. *In vitro* methodologies require significant inputs of materials and time, often at great cost.

5 C. Purine, Pyrimidine, Nucleoside and Nucleotide Metabolism and Uses

Purine and pyrimidine metabolism genes and their corresponding proteins are important targets for the therapy of tumor diseases and viral infections. The language "purine" or "pyrimidine" includes the nitrogenous bases which are constituents of nucleic acids, co-enzymes, and nucleotides. The term "nucleotide" includes the basic structural units of nucleic acid molecules, which are comprised of a nitrogenous base, a pentose sugar (in the case of RNA, the sugar is ribose; in the case of DNA, the sugar is D-deoxyribose), and phosphoric acid. The language "nucleoside" includes molecules which serve as precursors to nucleotides, but which are lacking the phosphoric acid moiety that nucleotides possess. By inhibiting the biosynthesis of these molecules, or their mobilization to form nucleic acid molecules, it is possible to inhibit RNA and DNA synthesis; by inhibiting this activity in a fashion targeted to cancerous cells, the ability of tumor cells to divide and replicate may be inhibited. Additionally, there are nucleotides which do not form nucleic acid molecules, but rather serve as energy stores (i.e., AMP) or as coenzymes (i.e., FAD and NAD).

Several publications have described the use of these chemicals for these medical indications, by influencing purine and/or pyrimidine metabolism (e.g. Christopherson, R.I. and Lyons, S.D. (1990) "Potent inhibitors of *de novo* pyrimidine and purine biosynthesis as chemotherapeutic agents." *Med. Res. Reviews* 10: 505-548). Studies of enzymes involved in purine and pyrimidine metabolism have been focused on the development of new drugs which can be used, for example, as immunosuppressants or anti-proliferants (Smith, J.L., (1995) "Enzymes in nucleotide synthesis." *Curr. Opin. Struct. Biol.* 5: 752-757; (1995) *Biochem Soc. Transact.* 23: 877-902). However, purine and pyrimidine bases, nucleosides and nucleotides have other utilities: as intermediates in the biosynthesis of several fine chemicals (e.g., thiamine, S-adenosyl-methionine, folates, or riboflavin), as energy carriers for the cell (e.g., ATP or GTP), and for chemicals themselves, commonly used as flavor enhancers (e.g., IMP or GMP) or for several medicinal applications (see, for example, Kuninaka, A. (1996) Nucleotides and

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Related Compounds in Biotechnology vol. 6, Rehm *et al.*, eds. VCH: Weinheim, p. 561-612). Also, enzymes involved in purine, pyrimidine, nucleoside, or nucleotide metabolism are increasingly serving as targets against which chemicals for crop protection, including fungicides, herbicides and insecticides, are developed.

The metabolism of these compounds in bacteria has been characterized (for 5 reviews see, for example, Zalkin, H. and Dixon, J.E. (1992) "de novo purine nucleotide biosynthesis", in: Progress in Nucleic Acid Research and Molecular Biology, vol. 42, Academic Press:, p. 259-287; and Michal, G. (1999) "Nucleotides and Nucleosides", Chapter 8 in: Biochemical Pathways: An Atlas of Biochemistry and Molecular Biology, Wiley: New York). Purine metabolism has been the subject of intensive research, and is 10 essential to the normal functioning of the cell. Impaired purine metabolism in higher animals can cause severe disease, such as gout. Purine nucleotides are synthesized from ribose-5-phosphate, in a series of steps through the intermediate compound inosine-5'phosphate (IMP), resulting in the production of guanosine-5'-monophosphate (GMP) or adenosine-5'-monophosphate (AMP), from which the triphosphate forms utilized as 15 nucleotides are readily formed. These compounds are also utilized as energy stores, so their degradation provides energy for many different biochemical processes in the cell. Pyrimidine biosynthesis proceeds by the formation of uridine-5'-monophosphate (UMP) from ribose-5-phosphate. UMP, in turn, is converted to cytidine-5'-triphosphate (CTP). The deoxy- forms of all of these nucleotides are produced in a one step reduction 20 reaction from the diphosphate ribose form of the nucleotide to the diphosphate deoxyribose form of the nucleotide. Upon phosphorylation, these molecules are able to participate in DNA synthesis.

25 D. Trehalose Metabolism and Uses

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Trehalose consists of two glucose molecules, bound in α, α-1,1 linkage. It is commonly used in the food industry as a sweetener, an additive for dried or frozen foods, and in beverages. However, it also has applications in the pharmaceutical, cosmetics and biotechnology industries (see, for example, Nishimoto *et al.*, (1998) U.S. Patent No. 5,759,610; Singer, M.A. and Lindquist, S. (1998) *Trends Biotech*. 16: 460-467; Paiva, C.L.A. and Panek, A.D. (1996) *Biotech. Ann. Rev.* 2: 293-314; and Shiosaka, M. (1997) J. Japan 172: 97-102). Trehalose is produced by enzymes from

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many microorganisms and is naturally released into the surrounding medium, from which it can be collected using methods known in the art.

II. Elements and Methods of the Invention

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The present invention is based, at least in part, on the discovery of novel molecules, referred to herein as MP nucleic acid and protein molecules, which play a role in or function in one or more cellular metabolic pathways. In one embodiment, the MP molecules catalyze an enzymatic reaction involving one or more amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathways. In a preferred embodiment, the activity of the MP molecules of the present invention in one or more *C. glutamicum* metabolic pathways for amino acids, vitamins, cofactors, nutraceuticals, nucleotides, nucleosides or trehalose has an impact on the production of a desired fine chemical by this organism. In a particularly preferred embodiment, the MP molecules of the invention are modulated in activity, such that the *C. glutamicum* metabolic pathways in which the MP proteins of the invention are involved are modulated in efficiency or output, which either directly or indirectly modulates the production or efficiency of production of a desired fine chemical by *C. glutamicum*.

The language, "MP protein" or "MP polypeptide" includes proteins which play a role in, *e.g.*, catalyze an enzymatic reaction, in one or more amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside or trehalose metabolic pathways. Examples of MP proteins include those encoded by the MP genes set forth in Table 1 and by the odd-numbered 3EQ 1D NOs. The terms MP gene or MP nucleic acid sequence" include nucleic acid sequences encoding an MP protein, which consist of a coding region and also corresponding untranslated 5' and 3' sequence regions. Examples of MP genes include those set forth in Table 1. The terms "production" or "productivity" are art-recognized and include the concentration of the fermentation product (for example, the desired fine chemical) formed within a given time and a given fermentation volume (*e.g.*, kg product per hour per liter). The term "efficiency of production" includes the time required for a particular level of production to be achieved (for example, how long it takes for the cell to attain a particular rate of output of a fine chemical). The term "yield" or "product/carbon yield" is art-recognized and includes

the efficiency of the conversion of the carbon source into the product (i.e., fine chemical). This is generally written as, for example, kg product per kg carbon source. By increasing the yield or production of the compound, the quantity of recovered molecules, or of useful recovered molecules of that compound in a given amount of culture over a given amount of time is increased. The terms "biosynthesis" or a "biosynthetic pathway" are art-recognized and include the synthesis of a compound, preferably an organic compound, by a cell from intermediate compounds in what may be a multistep and highly regulated process. The terms "degradation" or a "degradation pathway" are art-recognized and include the breakdown of a compound, preferably an organic compound, by a cell to degradation products (generally speaking, smaller or less complex molecules) in what may be a multistep and highly regulated process. The language "metabolism" is art-recognized and includes the totality of the biochemical reactions that take place in an organism. The metabolism of a particular compound, then, (e.g., the metabolism of an amino acid such as glycine) comprises the overall biosynthetic, modification, and degradation pathways in the cell related to this compound.

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In another embodiment, the MP molecules of the invention are capable of modulating the production of a desired molecule, such as a fine chemical, in a microorganism such as *C. glutamicum*. Using recombinant genetic techniques, one or more of the biosynthetic or degradative enzymes of the invention for amino acids, vitamins, cofactors, nutraceuticals, nucleotides, nucleosides, or trehalose may be manipulated such that its function is modulated. For example, a biosynthetic enzyme may be improved in efficiency, or its allosteric control region destroyed such that feedback inhibition of production of the compound is prevented. Similarly, a degradative enzyme may be deleted or modified by substitution, deletion, or addition such that its degradative activity is lessened for the desired compound without impairing the viability of the cell. In each case, the overall yield or rate of production of one of these desired fine chemicals may be increased.

It is also possible that such alterations in the protein and nucleotide molecules of the invention may improve the production of other fine chemicals besides the amino acids, vitamins, cofactors, nutraceuticals, nucleotides, nucleosides, and trehalose.

Metabolism of any one compound is necessarily intertwined with other biosynthetic and

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degradative pathways within the cell, and necessary cofactors, intermediates, or substrates in one pathway are likely supplied or limited by another such pathway. Therefore, by modulating the activity of one or more of the proteins of the invention, the production or efficiency of activity of another fine chemical biosynthetic or degradative pathway may be impacted. For example, amino acids serve as the structural units of all proteins, yet may be present intracellularly in levels which are limiting for protein synthesis; therefore, by increasing the efficiency of production or the yields of one or more amino acids within the cell, proteins, such as biosynthetic or degradative proteins, may be more readily synthesized. Likewise, an alteration in a metabolic pathway enzyme such that a particular side reaction becomes more or less favored may result in the over- or under-production of one or more compounds which are utilized as intermediates or substrates for the production of a desired fine chemical.

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The isolated nucleic acid sequences of the invention are contained within the genome of a *Corynebacterium glutamicum* strain available through the American Type Culture Collection, given designation ATCC 13032. The nucleotide sequence of the isolated *C. glutamicum* MP DNAs and the predicted amino acid sequences of the *C. glutamicum* MP proteins are shown in the Sequence Listing as odd-numbered SEQ ID NOs and even-numbered SEQ ID NOs, respectively. Computational analyses were performed which classified and/or identified these nucleotide sequences as sequences which encode metabolic pathway proteins.

The present invention also pertains to proteins which have an amino acid sequence which is substantially homologous to an amino acid sequence of the invention (e.g., the sequence of an even-numbered SEQ ID NO of the Sequence Listing). As used herein, a protein which has an amino acid sequence which is substantially homologous to a selected amino acid sequence is least about 50% homologous to the selected amino acid sequence, e.g., the entire selected amino acid sequence. A protein which has an amino acid sequence which is substantially homologous to a selected amino acid sequence can also be least about 50-60%, preferably at least about 60-70%, and more preferably at least about 70-80%, 80-90%, or 90-95%, and most preferably at least about 96%, 97%, 98%, 99% or more homologous to the selected amino acid sequence.

The MP protein or a biologically active portion or fragment thereof of the invention can catalyze an enzymatic reaction in one or more amino acid, vitamin,

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cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathways, or have one or more of the activities set forth in Table 1.

Various aspects of the invention are described in further detail in the following subsections:

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A. Isolated Nucleic Acid Molecules

One aspect of the invention pertains to isolated nucleic acid molecules that encode MP polypeptides or biologically active portions thereof, as well as nucleic acid fragments sufficient for use as hybridization probes or primers for the identification or amplification of MP-encoding nucleic acid (e.g., MP DNA). As used herein, the term "nucleic acid molecule" is intended to include DNA molecules (e.g., cDNA or genomic DNA) and RNA molecules (e.g., mRNA) and analogs of the DNA or RNA generated using nucleotide analogs. This term also encompasses untranslated sequence located at both the 3' and 5' ends of the coding region of the gene: at least about 100 nucleotides of sequence upstream from the 5' end of the coding region and at least about 20 nucleotides of sequence downstream from the 3'end of the coding region of the gene. The nucleic acid molecule can be single-stranded or double-stranded, but preferably is double-stranded DNA. An "isolated" nucleic acid molecule is one which is separated from other nucleic acid molecules which are present in the natural source of the nucleic acid. Preferably, an "isolated" nucleic acid is free of sequences which naturally flank the nucleic acid (i.e., sequences located at the 5' and 3' ends of the nucleic acid) in the genomic DNA of the organism from which the nucleic acid is derived. For example, in various embodiments, the isolated MP nucleic acid molecule can contain less than about 5 kb, 4kb, 3kb, 2kb, 1 kb, 0.5 kb or 0.1 kb of nucleotide sequences which naturally flank the nucleic acid molecule in genomic DNA of the cell from which the nucleic acid is derived (e.g., a C. glutamicum cell). Moreover, an "isolated" nucleic acid molecule, such as a DNA molecule, can be substantially free of other cellular material, or culture medium when produced by recombinant techniques, or chemical precursors or other chemicals when chemically synthesized.

A nucleic acid molecule of the present invention, *e.g.*, a nucleic acid molecule having a nucleotide sequence of an odd-numbered SEQ ID NO of the Sequence Listing, or a portion thereof, can be isolated using standard molecular biology techniques and the

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sequence information provided herein. For example, a C. glutamicum MP DNA can be isolated from a C. glutamicum library using all or portion of one of the odd-numbered SEQ ID NO sequences of the Sequence Listing as a hybridization probe and standard hybridization techniques (e.g., as described in Sambrook, J., Fritsh, E. F., and Maniatis, T. Molecular Cloning: A Laboratory Manual. 2nd, ed., Cold Spring Harbor Laboratory, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989). Moreover, a nucleic acid molecule encompassing all or a portion of one of the nucleic acid sequences of the invention (e.g., an odd-numbered SEQ ID NO:) can be isolated by the polymerase chain reaction using oligonucleotide primers designed based upon this sequence (e.g., a nucleic acid molecule encompassing all or a portion of one of the 10 nucleic acid sequences of the invention (e.g., an odd-numbered SEQ ID NO of the Sequence Listing) can be isolated by the polymerase chain reaction using oligonucleotide primers designed based upon this same sequence). For example, mRNA can be isolated from normal endothelial cells (e.g., by the guanidinium-thiocyanate extraction procedure of Chirgwin et al. (1979) Biochemistry 18: 5294-5299) and DNA 15 can be prepared using reverse transcriptase (e.g., Moloney MLV reverse transcriptase, available from Gibco/BRL, Bethesda, MD; or AMV reverse transcriptase, available from Seikagaku America, Inc., St. Petersburg, FL). Synthetic oligonucleotide primers for polymerase chain reaction amplification can be designed based upon one of the 20 nucleotide sequences shown in the Sequence Listing. A nucleic acid of the invention can be amplified using cDNA or, alternatively, genomic DNA, as a template and appropriate oligonucleotide primers according to standard PCR amplification techniques. The nucleic acid so amplified can be cloned into an appropriate vector and characterized by DNA sequence analysis. Furthermore, oligonucleotides corresponding

In a preferred embodiment, an isolated nucleic acid molecule of the invention comprises one of the nucleotide sequences shown in the Sequence Listing. The nucleic acid sequences of the invention, as set forth in the Sequence Listing, correspond to the *Corynebacterium glutamicum* MP DNAs of the invention. This DNA comprises sequences encoding MP proteins (*i.e.*, the "coding region", indicated in each odd-numbered SEQ ID NO: sequence in the Sequence Listing), as well as 5' untranslated

to an MP nucleotide sequence can be prepared by standard synthetic techniques, e.g.,

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using an automated DNA synthesizer.

sequences and 3' untranslated sequences, also indicated in each odd-numbered SEQ ID NO: in the Sequence Listing. Alternatively, the nucleic acid molecule can comprise only the coding region of any of the nucleic acid sequences of the Sequence Listing.

For the purposes of this application, it will be understood that each of the nucleic acid and amino acid sequences set forth in the Sequence Listing has an identifying RXA, 5 RXN, RXS, or RXC number having the designation "RXA", "RXN", "RXS", or "RXC" followed by 5 digits (i.e., RXA00007, RXN00023, RXS00116, or RXC00128). Each of the nucleic acid sequences comprises up to three parts: a 5' upstream region, a coding region, and a downstream region. Each of these three regions is identified by the same RXA, RXN, RXS, or RXC designation to eliminate confusion. The recitation "one of the odd-numbered sequences of the Sequence Listing", then, refers to any of the nucleic acid sequences in the Sequence Listing, which may also be distinguished by their differing RXA, RXN, RXS, or RXC designations. The coding region of each of these sequences is translated into a corresponding amino acid sequence, which is also set forth in the Sequence Listing, as an even-numbered SEQ ID NO: immediately following the corresponding nucleic acid sequence . For example, the coding region for RXA02229 is set forth in SEQ ID NO:1, while the amino acid sequence which it encodes is set forth as SEQ ID NO:2. The sequences of the nucleic acid molecules of the invention are identified by the same RXA, RXN, RXS, or RXC designations as the amino acid molecules which they encode, such that they can be readily correlated. For example, the 20 amino acid sequences designated RXA02229, RX00351, RXS02970, and RXC02390 are translations of the coding regions of the nucleotide sequences of nucleic acid molecules RXA02229, RX00351, RXS02970, and RXC02390, respectively. The correspondence between the RXA, RXN, RXS, and RXC nucleotide and amino acid sequences of the invention and their assigned SEQ ID NOs is set forth in Table 1. 25

Several of the genes of the invention are "F-designated genes". An F-designated gene includes those genes set forth in Table 1 which have an 'F' in front of the RXA, RXN, RXS, or RXC designation. For example, SEQ ID NO:5, designated, as indicated on Table 1, as "F RXA01009", is an F-designated gene, as are SEQ ID NOs: 73, 75, and 77 (designated on Table 1 as "F RXA00007", "F RXA00364", and "F RXA00367", respectively).

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In one embodiment, the nucleic acid molecules of the present invention are not intended to include *C. glutamicum* those compiled in Table 2. In the case of the dapD gene, a sequence for this gene was published in Wehrmann, A., *et al.* (1998) *J. Bacteriol.* 180(12): 3159-3165. However, the sequence obtained by the inventors of the present application is significantly longer than the published version. It is believed that the published version relied on an incorrect start codon, and thus represents only a fragment of the actual coding region.

In another preferred embodiment, an isolated nucleic acid molecule of the invention comprises a nucleic acid molecule which is a complement of one of the nucleotide sequences of the invention (e.g., a sequence of an odd-numbered SEQ ID NO: of the Sequence Listing), or a portion thereof. A nucleic acid molecule which is complementary to one of the nucleotide sequences of the invention is one which is sufficiently complementary to one of the nucleotide sequences shown in the Sequence Listing (e.g., the sequence of an odd-numbered SEQ ID NO:) such that it can hybridize to one of the nucleotide sequences of the invention, thereby forming a stable duplex.

In still another preferred embodiment, an isolated nucleic acid molecule of the invention comprises a nucleotide sequence which is at least about 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, or 60%, preferably at least about 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, or 70%%, more preferably at least about 71%, 72%, 73%, 74%, 75%, 76%, 77%, 78%, 79%, or 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, or 90%, or 91%, 92%, 93%, 94%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more homologous to a nucleotide sequence of the invention (e.g., a sequence of an odd-numbered SEQ ID NO: of the Sequence Listing), or a portion thereof. Ranges and identity values intermediate to the above-recited ranges, (e.g., 70-90% identical or 80-95% identical) are also intended to be encompassed by the present invention. For example, ranges of identity values using a combination of any of the above values recited as upper and/or lower limits are intended to be included. In an additional preferred embodiment, an isolated nucleic acid molecule of the invention comprises a nucleotide sequence which hybridizes, e.g., hybridizes under stringent conditions, to one of the nucleotide sequences of the invention, or a portion thereof.

Moreover, the nucleic acid molecule of the invention can comprise only a portion of the coding region of the sequence of one of the odd-numbered SEQ ID NOs

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of the Sequence Listing, for example a fragment which can be used as a probe or primer or a fragment encoding a biologically active portion of an MP protein. The nucleotide sequences determined from the cloning of the MP genes from C. glutamicum allows for the generation of probes and primers designed for use in identifying and/or cloning MP homologues in other cell types and organisms, as well as MP homologues from other Corynebacteria or related species. The probe/primer typically comprises substantially purified oligonucleotide. The oligonucleotide typically comprises a region of nucleotide sequence that hybridizes under stringent conditions to at least about 12, preferably about 25, more preferably about 40, 50 or 75 consecutive nucleotides of a sense strand of one of the nucleotide sequences of the invention (e.g., a sequence of one of the oddnumbered SEQ ID NOs of the Sequence Listing), an anti-sense sequence of one of these sequences, or naturally occurring mutants thereof. Primers based on a nucleotide sequence of the invention can be used in PCR reactions to clone MP homologues. Probes based on the MP nucleotide sequences can be used to detect transcripts or genomic sequences encoding the same or homologous proteins. In preferred embodiments, the probe further comprises a label group attached thereto, e.g. the label group can be a radioisotope, a fluorescent compound, an enzyme, or an enzyme cofactor. Such probes can be used as a part of a diagnostic test kit for identifying cells which misexpress an MP protein, such as by measuring a level of an MP-encoding nucleic acid in a sample of cells from a subject e.g., detecting MP mRNA levels or determining whether a genomic MP gene has been mutated or deleted.

In one embodiment, the nucleic acid molecule of the invention encodes a protein or portion thereof which includes an amino acid sequence which is sufficiently homologous to an amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO of the Sequence Listing) such that the protein or portion thereof maintains the ability to catalyze an enzymatic reaction in an amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathway. As used herein, the language "sufficiently homologous" refers to proteins or portions thereof which have amino acid sequences which include a minimum number of identical or equivalent (e.g., an amino acid residue which has a similar side chain as an amino acid residue in a sequence of one of the even-numbered SEQ ID NOs of the Sequence Listing) amino acid residues to an amino acid sequence of the invention such that the

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protein or portion thereof is able to catalyze an enzymatic reaction in a *C. glutamicum* amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside or trehalose metabolic pathway. Protein members of such metabolic pathways, as described herein, function to catalyze the biosynthesis or degradation of one or more of: amino acids, vitamins, cofactors, nutraceuticals, nucleotides, nucleosides, or trehalose. Examples of such activities are also described herein. Thus, "the function of an MP protein" contributes to the overall functioning of one or more such metabolic pathway and contributes, either directly or indirectly, to the yield, production, and/or efficiency of production of one or more fine chemicals. Examples of MP protein activities are set

In another embodiment, the protein is at least about 50-60%, preferably at least about 60-70%, and more preferably at least about 70-80%, 80-90%, 90-95%, and most preferably at least about 96%, 97%, 98%, 99% or more homologous to an entire amino acid sequence of the invention (*e.g.*, a sequence of an even-numbered SEQ ID NO: of the Sequence Listing).

Portions of proteins encoded by the MP nucleic acid molecules of the invention are preferably biologically active portions of one of the MP proteins. As used herein, the term "biologically active portion of an MP protein" is intended to include a portion, e.g., a domain/motif, of an MP protein that catalyzes an enzymatic reaction in one or more C. glutamicum amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathways, or has an activity as set forth in Table 1. To determine whether an MP protein or a biologically active portion thereof can catalyze an enzymatic reaction in an amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathway, an assay of enzymatic activity may be performed. Such assay methods are well known to those of ordinary skill in the art, as detailed in Example 8 of the Exemplification.

Additional nucleic acid fragments encoding biologically active portions of an MP protein can be prepared by isolating a portion of one of the amino acid sequences of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing), expressing the encoded portion of the MP protein or peptide (e.g., by recombinant expression *in vitro*) and assessing the activity of the encoded portion of the MP protein or peptide.

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The invention further encompasses nucleic acid molecules that differ from one of the nucleotide sequences of the invention (e.g., a sequence of an odd-numbered SEQ ID NO: of the Sequence Listing) (and portions thereof) due to degeneracy of the genetic code and thus encode the same MP protein as that encoded by the nucleotide sequences of the invention. In another embodiment, an isolated nucleic acid molecule of the invention has a nucleotide sequence encoding a protein having an amino acid sequence shown in the Sequence Listing (e.g., an even-numbered SEQ ID NO:). In a still further embodiment, the nucleic acid molecule of the invention encodes a full length *C*. *glutamicum* protein which is substantially homologous to an amino acid sequence of the invention (encoded by an open reading frame shown in an odd-numbered SEQ ID NO: of the Sequence Listing).

It will be understood by one of ordinary skill in the art that in one embodiment the sequences of the invention are not meant to include the sequences of the prior art, such as those Genbank sequences set forth in Tables 2 or 4 which were available prior to the present invention. In one embodiment, the invention includes nucleotide and amino acid sequences having a percent identity to a nucleotide or amino acid sequence of the invention which is greater than that of a sequence of the prior art (e.g., a Genbank sequence (or the protein encoded by such a sequence) set forth in Tables 2 or 4). For example, the invention includes a nucleotide sequence which is greater than and/or at least 40% identical to the nucleotide sequence designated RXA00115 (SEQ ID NO:185), a nucleotide sequence which is greater than and/or at least % identical to the nucleotide sequence designated RXA00131 (SEQ ID NO:991), and a nucleotide sequence which is greater than and/or at least 39% identical to the nucleotide sequence designated RXA00219 (SEQ ID NO:345). One of ordinary skill in the art would be able to calculate the lower threshold of percent identity for any given sequence of the invention by examining the GAP-calculated percent identity scores set forth in Table 4 for each of the three top hits for the given sequence, and by subtracting the highest GAP-calculated percent identity from 100 percent. One of ordinary skill in the art will also appreciate that nucleic acid and amino acid sequences having percent identities greater than the lower threshold so calculated (e.g., at least 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, or 60%, preferably at least about 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, or 70%, more preferably at least about 71%, 72%, 73%,

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74%, 75%, 76%, 77%, 78%, 79%, or 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, or 90%, or 91%, 92%, 93%, 94%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more identical) are also encompassed by the invention.

In addition to the *C. glutamicum* MP nucleotide sequences set forth in the Sequence Listing as odd-numbered SEQ ID NOs, it will be appreciated by one of ordinary skill in the art that DNA sequence polymorphisms that lead to changes in the amino acid sequences of MP proteins may exist within a population (*e.g.*, the *C. glutamicum* population). Such genetic polymorphism in the MP gene may exist among individuals within a population due to natural variation. As used herein, the terms "gene" and "recombinant gene" refer to nucleic acid molecules comprising an open reading frame encoding an MP protein, preferably a *C. glutamicum* MP protein. Such natural variations can typically result in 1-5% variance in the nucleotide sequence of the MP gene. Any and all such nucleotide variations and resulting amino acid polymorphisms in MP that are the result of natural variation and that do not alter the functional activity of MP proteins are intended to be within the scope of the invention.

Nucleic acid molecules corresponding to natural variants and non-C. glutamicum homologues of the C. glutamicum MP DNA of the invention can be isolated based on their homology to the C. glutamicum MP nucleic acid disclosed herein using the C. glutamicum DNA, or a portion thereof, as a hybridization probe according to standard hybridization techniques under stringent hybridization conditions. Accordingly, in another embodiment, an isolated nucleic acid molecule of the invention is at least 15 nucleotides in length and hybridizes under stringent conditions to the nucleic acid molecule comprising a nucleotide sequence of an odd-numbered SEQ ID NO: of the Sequence Listing. In other embodiments, the nucleic acid is at least 30, 50, 100, 250 or more nucleotides in length. As used herein, the term "hybridizes under stringent conditions" is intended to describe conditions for hybridization and washing under which nucleotide sequences at least 60% homologous to each other typically remain hybridized to each other. Preferably, the conditions are such that sequences at least about 65%, more preferably at least about 70%, and even more preferably at least about 75% or more homologous to each other typically remain hybridized to each other. Such stringent conditions are known to one of ordinary skill in the art and can be found in Current Protocols in Molecular Biology, John Wiley & Sons, N.Y. (1989), 6.3.1-6.3.6.

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A preferred, non-limiting example of stringent hybridization conditions are hybridization in 6X sodium chloride/sodium citrate (SSC) at about 45°C, followed by one or more washes in 0.2 X SSC, 0.1% SDS at 50-65°C. Preferably, an isolated nucleic acid molecule of the invention that hybridizes under stringent conditions to a nucleotide sequence of the invention corresponds to a naturally-occurring nucleic acid molecule. As used herein, a "naturally-occurring" nucleic acid molecule refers to an RNA or DNA molecule having a nucleotide sequence that occurs in nature (e.g., encodes a natural protein). In one embodiment, the nucleic acid encodes a natural *C. glutamicum* MP protein.

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In addition to naturally-occurring variants of the MP sequence that may exist in the population, one of ordinary skill in the art will further appreciate that changes can be introduced by mutation into a nucleotide sequence of the invention, thereby leading to changes in the amino acid sequence of the encoded MP protein, without altering the functional ability of the MP protein. For example, nucleotide substitutions leading to amino acid substitutions at "non-essential" amino acid residues can be made in a nucleotide sequence of the invention. A "non-essential" amino acid residue is a residue that can be altered from the wild-type sequence of one of the MP proteins (e.g., an even-numbered SEQ ID NO: of the Sequence Listing) without altering the activity of said MP protein, whereas an "essential" amino acid residue is required for MP protein activity. Other amino acid residues, however, (e.g., those that are not conserved or only semi-conserved in the domain having MP activity) may not be essential for activity and thus are likely to be amenable to alteration without altering MP activity.

Accordingly, another aspect of the invention pertains to nucleic acid molecules encoding MP proteins that contain changes in amino acid residues that are not essential for MP activity. Such MP proteins differ in amino acid sequence from a sequence of an even-numbered SEQ ID NO: of the Sequence Listing yet retain at least one of the MP activities described herein. In one embodiment, the isolated nucleic acid molecule comprises a nucleotide sequence encoding a protein, wherein the protein comprises an amino acid sequence at least about 50% homologous to an amino acid sequence of the invention and is capable of catalyzing an enzymatic reaction in an amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathway, or has one or more activities set forth in Table 1. Preferably, the protein encoded by the nucleic

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acid molecule is at least about 50-60% homologous to the amino acid sequence of one of the odd-numbered SEQ ID NOs of the Sequence Listing, more preferably at least about 60-70% homologous to one of these sequences, even more preferably at least about 70-80%, 80-90%, 90-95% homologous to one of these sequences, and most preferably at least about 96%, 97%, 98%, or 99% homologous to one of the amino acid sequences of the invention.

To determine the percent homology of two amino acid sequences (*e.g.*, one of the amino acid sequences of the invention and a mutant form thereof) or of two nucleic acids, the sequences are aligned for optimal comparison purposes (*e.g.*, gaps can be introduced in the sequence of one protein or nucleic acid for optimal alignment with the other protein or nucleic acid). The amino acid residues or nucleotides at corresponding amino acid positions or nucleotide positions are then compared. When a position in one sequence (*e.g.*, one of the amino acid sequences of the invention) is occupied by the same amino acid residue or nucleotide as the corresponding position in the other sequence (*e.g.*, a mutant form of the amino acid sequence), then the molecules are homologous at that position (*i.e.*, as used herein amino acid or nucleic acid "homology" is equivalent to amino acid or nucleic acid "identity"). The percent homology between the two sequences is a function of the number of identical positions shared by the sequences (*i.e.*, % homology = # of identical positions/total # of positions x 100).

An isolated nucleic acid molecule encoding an MP protein homologous to a protein sequence of the invention (*e.g.*, a sequence of an even-numbered SEQ ID NO: of the Sequence Listing) can be created by introducing one or more nucleotide substitutions, additions or deletions into a nucleotide sequence of the invention such that one or more amino acid substitutions, additions or deletions are introduced into the encoded protein. Mutations can be introduced into one of the nucleotide sequences of the invention by standard techniques, such as site-directed mutagenesis and PCR-mediated mutagenesis. Preferably, conservative amino acid substitutions are made at one or more predicted non-essential amino acid residues. A "conservative amino acid substitution" is one in which the amino acid residue is replaced with an amino acid residue having a similar side chain. Families of amino acid residues having similar side chains have been defined in the art. These families include amino acids with basic side chains (*e.g.*, lysine, arginine, histidine), acidic side chains (*e.g.*, aspartic acid. glutamic

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acid), uncharged polar side chains (*e.g.*, glycine, asparagine, glutamine, serine, threonine, tyrosine, cysteine), nonpolar side chains (*e.g.*, alanine, valine, leucine, isoleucine, proline, phenylalanine, methionine, tryptophan), beta-branched side chains (*e.g.*, threonine, valine, isoleucine) and aromatic side chains (*e.g.*, tyrosine, phenylalanine, tryptophan, histidine). Thus, a predicted nonessential amino acid residue in an MP protein is preferably replaced with another amino acid residue from the same side chain family. Alternatively, in another embodiment, mutations can be introduced randomly along all or part of an MP coding sequence, such as by saturation mutagenesis, and the resultant mutants can be screened for an MP activity described herein to identify mutants that retain MP activity. Following mutagenesis of the nucleotide sequence of one of the odd-numbered SEQ ID NOs of the Sequence Listing, the encoded protein can be expressed recombinantly and the activity of the protein can be determined using, for example, assays described herein (see Example 8 of the Exemplification).

In addition to the nucleic acid molecules encoding MP proteins described above, another aspect of the invention pertains to isolated nucleic acid molecules which are antisense thereto. An "antisense" nucleic acid comprises a nucleotide sequence which is complementary to a "sense" nucleic acid encoding a protein, e.g., complementary to the coding strand of a double-stranded DNA molecule or complementary to an mRNA sequence. Accordingly, an antisense nucleic acid can hydrogen bond to a sense nucleic acid. The antisense nucleic acid can be complementary to an entire MP coding strand, or to only a portion thereof. In one embodiment, an antisense nucleic acid molecule is antisense to a "coding region" of the coding strand of a nucleotide sequence encoding an MP protein. The term "coding region" refers to the region of the nucleotide sequence comprising codons which are translated into amino acid residues (e.g., the entire coding region of SEQ ID NO. 1 (RXA02229) comprises nucleotides 1 to 825). In another embodiment, the antisense nucleic acid molecule is antisense to a "noncoding region" of the coding strand of a nucleotide sequence encoding MP. The term "noncoding region" refers to 5' and 3' sequences which flank the coding region that are not translated into amino acids (i.e., also referred to as 5' and 3' untranslated regions).

Given the coding strand sequences encoding MP disclosed herein (e.g., the sequences set forth as odd-numbered SEQ ID NOs in the Sequence Listing), antisense

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nucleic acids of the invention can be designed according to the rules of Watson and Crick base pairing. The antisense nucleic acid molecule can be complementary to the entire coding region of MP mRNA, but more preferably is an oligonucleotide which is antisense to only a portion of the coding or noncoding region of MP mRNA. For example, the antisense oligonucleotide can be complementary to the region surrounding the translation start site of MP mRNA. An antisense oligonucleotide can be, for example, about 5, 10, 15, 20, 25, 30, 35, 40, 45 or 50 nucleotides in length. An antisense nucleic acid of the invention can be constructed using chemical synthesis and enzymatic ligation reactions using procedures known in the art. For example, an antisense nucleic acid (e.g., an antisense oligonucleotide) can be chemically synthesized 10 using naturally occurring nucleotides or variously modified nucleotides designed to increase the biological stability of the molecules or to increase the physical stability of the duplex formed between the antisense and sense nucleic acids, e.g., phosphorothioate derivatives and acridine substituted nucleotides can be used. Examples of modified nucleotides which can be used to generate the antisense nucleic acid include 5-15 fluorouracil, 5-bromouracil, 5-chlorouracil, 5-iodouracil, hypoxanthine, xanthine, 4acetylcytosine, 5-(carboxyhydroxylmethyl) uracil, 5-carboxymethylaminomethyl-2thiouridine, 5-carboxymethylaminomethyluracil, dihydrouracil, beta-Dgalactosylqueosine, inosine, N6-isopentenyladenine, 1-methylguanine, 1-methylinosine, 20 2,2-dimethylguanine, 2-methyladenine, 2-methylguanine, 3-methylcytosine, 5methylcytosine, N6-adenine, 7-methylguanine, 5-methylaminomethyluracil, 5methoxyaminomethyl-2-thiouracil, beta-D-mannosylqueosine, 5'methoxyearboxymethyluraeil, 5-methoxyuraeil, 2-methyltnio-No-isopentenyladenine, uracil-5-oxyacetic acid (v), wybutoxosine, pseudouracil, queosine, 2-thiocytosine, 5methyl-2-thiouracil, 2-thiouracil, 4-thiouracil, 5-methyluracil, uracil-5- oxyacetic acid 25 methylester, uracil-5-oxyacetic acid (v), 5-methyl-2-thiouracil, 3-(3-amino-3-N-2carboxypropyl) uracil, (acp3)w, and 2,6-diaminopurine. Alternatively, the antisense nucleic acid can be produced biologically using an expression vector into which a nucleic acid has been subcloned in an antisense orientation (i.e., RNA transcribed from the inserted nucleic acid will be of an antisense orientation to a target nucleic acid of 30 interest, described further in the following subsection).

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The antisense nucleic acid molecules of the invention are typically administered to a cell or generated *in situ* such that they hybridize with or bind to cellular mRNA and/or genomic DNA encoding an MCT protein to thereby inhibit expression of the protein, *e.g.*, by inhibiting transcription and/or translation. The hybridization can be by conventional nucleotide complementarity to form a stable duplex, or, for example, in the case of an antisense nucleic acid molecule which binds to DNA duplexes, through specific interactions in the major groove of the double helix. The antisense molecule can be modified such that it specifically binds to a receptor or an antigen expressed on a selected cell surface, *e.g.*, by linking the antisense nucleic acid molecule to a peptide or an antibody which binds to a cell surface receptor or antigen. The antisense nucleic acid molecule can also be delivered to cells using the vectors described herein. To achieve sufficient intracellular concentrations of the antisense molecules, vector constructs in which the antisense nucleic acid molecule is placed under the control of a strong prokaryotic, viral, or eukaryotic promoter are preferred.

In yet another embodiment, the antisense nucleic acid molecule of the invention is an α-anomeric nucleic acid molecule. An α-anomeric nucleic acid molecule forms specific double-stranded hybrids with complementary RNA in which, contrary to the usual β-units, the strands run parallel to each other (Gaultier *et al.* (1987) *Nucleic Acids*. *Res.* 15:6625-6641). The antisense nucleic acid molecule can also comprise a 2'-o-methylribonucleotide (Inoue *et al.* (1987) *Nucleic Acids Res.* 15:6131-6148) or a chimeric RNA-DNA analogue (Inoue *et al.* (1987) *FEBS Lett.* 215:327-330).

In still another embodiment, an antisense nucleic acid of the invention is a ribozyme. Ribozymes are catalytic RNA molecules with ribonuclease activity which are capable of cleaving a single-stranded nucleic acid, such as an mRNA, to which they have a complementary region. Thus, ribozymes (e.g., hammerhead ribozymes (described in Haselhoff and Gerlach (1988) *Nature* 334:585-591)) can be used to catalytically cleave MP mRNA transcripts to thereby inhibit translation of MP mRNA. A ribozyme having specificity for an MP-encoding nucleic acid can be designed based upon the nucleotide sequence of an MP DNA disclosed herein (i.e., SEQ ID NO: 1 (RXA02229). For example, a derivative of a *Tetrahymena* L-19 IVS RNA can be constructed in which the nucleotide sequence of the active site is complementary to the nucleotide sequence to be cleaved in an MP-encoding mRNA. See, e.g., Cech et al.

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U.S. Patent No. 4,987,071 and Cech *et al.* U.S. Patent No. 5,116,742. Alternatively, MP mRNA can be used to select a catalytic RNA having a specific ribonuclease activity from a pool of RNA molecules. See, *e.g.*, Bartel, D. and Szostak, J.W. (1993) *Science* 261:1411-1418.

Alternatively, MP gene expression can be inhibited by targeting nucleotide sequences complementary to the regulatory region of an MP nucleotide sequence (e.g., an MP promoter and/or enhancers) to form triple helical structures that prevent transcription of an MP gene in target cells. See generally, Helene, C. (1991) Anticancer Drug Des. 6(6):569-84; Helene, C. et al. (1992) Ann. N.Y. Acad. Sci. 660:27-36; and Maher, L.J. (1992) Bioassays 14(12):807-15.

B. Recombinant Expression Vectors and Host Cells

Another aspect of the invention pertains to vectors, preferably expression vectors, containing a nucleic acid encoding an MP protein (or a portion thereof). As used herein, the term "vector" refers to a nucleic acid molecule capable of transporting another nucleic acid to which it has been linked. One type of vector is a "plasmid", which refers to a circular double stranded DNA loop into which additional DNA segments can be ligated. Another type of vector is a viral vector, wherein additional DNA segments can be ligated into the viral genome. Certain vectors are capable of autonomous replication in a host cell into which they are introduced (e.g., bacterial vectors having a bacterial origin of replication and episomal mammalian vectors). Other vectors (e.g., non-episomal mammalian vectors) are integrated into the genome of a host cell apon introduction into the host cell, and thereby are replicated along with the host genome. Moreover, certain vectors are capable of directing the expression of genes to which they are operatively linked. Such vectors are referred to herein as "expression vectors". In general, expression vectors of utility in recombinant DNA techniques are often in the form of plasmids. In the present specification, "plasmid" and "vector" can be used interchangeably as the plasmid is the most commonly used form of vector. However, the invention is intended to include such other forms of expression vectors, such as viral vectors (e.g., replication defective retroviruses, adenoviruses and adenoassociated viruses), which serve equivalent functions.

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The recombinant expression vectors of the invention comprise a nucleic acid of the invention in a form suitable for expression of the nucleic acid in a host cell, which means that the recombinant expression vectors include one or more regulatory sequences, selected on the basis of the host cells to be used for expression, which is operatively linked to the nucleic acid sequence to be expressed. Within a recombinant 5 expression vector, "operably linked" is intended to mean that the nucleotide sequence of interest is linked to the regulatory sequence(s) in a manner which allows for expression of the nucleotide sequence (e.g., in an in vitro transcription/translation system or in a host cell when the vector is introduced into the host cell). The term "regulatory sequence" is intended to include promoters, repressor binding sites, activator binding 10 sites, enhancers and other expression control elements (e.g., terminators, polyadenylation signals, or other elements of mRNA secondary structure). Such regulatory sequences are described, for example, in Goeddel; Gene Expression Technology: Methods in Enzymology 185, Academic Press, San Diego, CA (1990). Regulatory sequences include those which direct constitutive expression of a nucleotide 15 sequence in many types of host cell and those which direct expression of the nucleotide sequence only in certain host cells. Preferred regulatory sequences are, for example, promoters such as cos-, tac-, trp-, tet-, trp-tet-, lpp-, lac-, lpp-lac-, lac1^q-, T7-, T5-, T3-, gal-, trc-, ara-, SP6-, arny, SPO2, λ-P_R- or λ P_L, which are used preferably in bacteria. Additional regulatory sequences are, for example, promoters from yeasts and fungi, such 20 as ADC1, MFa, AC, P-60, CYC1, GAPDH, TEF, rp28, ADH, promoters from plants such as CaMV/35S, SSU, OCS, lib4, usp, STLS1, B33, nos or ubiquitin- or phaseolinpromoters. It is also possible to use artificial promoters. It will be appreciated by one of ordinary skill in the art that the design of the expression vector can depend on such factors as the choice of the host cell to be transformed, the level of expression of protein 25 desired, etc. The expression vectors of the invention can be introduced into host cells to thereby produce proteins or peptides, including fusion proteins or peptides, encoded by nucleic acids as described herein (e.g., MP proteins, mutant forms of MP proteins, fusion proteins, etc.).

The recombinant expression vectors of the invention can be designed for expression of MP proteins in prokaryotic or eukaryotic cells. For example, MP genes can be expressed in bacterial cells such as *C. glutamicum*, insect cells (using baculovirus

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"Foreign gene expression in yeast: a review", *Yeast* 8: 423-488; van den Hondel, C.A.M.J.J. *et al.* (1991) "Heterologous gene expression in filamentous fungi" in: More Gene Manipulations in Fungi, J.W. Bennet & L.L. Lasure, eds., p. 396-428: Academic Press: San Diego; and van den Hondel, C.A.M.J.J. & Punt, P.J. (1991) "Gene transfer systems and vector development for filamentous fungi, in: Applied Molecular Genetics of Fungi, Peberdy, J.F. *et al.*, eds., p. 1-28, Cambridge University Press: Cambridge), algae and multicellular plant cells (see Schmidt, R. and Willmitzer, L. (1988) High efficiency *Agrobacterium tumefaciens* –mediated transformation of *Arabidopsis thaliana* leaf and cotyledon explants" *Plant Cell Rep.*: 583-586), or mammalian cells. Suitable host cells are discussed further in Goeddel, *Gene Expression Technology: Methods in Enzymology* 185, Academic Press, San Diego, CA (1990). Alternatively, the recombinant expression vector can be transcribed and translated *in vitro*, for example using T7 promoter regulatory sequences and T7 polymerase.

Expression of proteins in prokaryotes is most often carried out with vectors containing constitutive or inducible promoters directing the expression of either fusion or non-fusion proteins. Fusion vectors add a number of amino acids to a protein encoded therein, usually to the amino terminus of the recombinant protein but also to the C-terminus or fused within suitable regions in the proteins. Such fusion vectors typically serve three purposes: 1) to increase expression of recombinant protein; 2) to increase the solubility of the recombinant protein; and 3) to aid in the purification of the recombinant protein by acting as a ligand in affinity purification. Often, in fusion expression vectors, a proteolytic cleavage site is introduced at the junction of the fusion moiety and the recombinant protein to enable separation of the recombinant protein from the fusion moiety subsequent to purification of the fusion protein. Such enzymes, and their cognate recognition sequences, include Factor Xa, thrombin and enterokinase.

Typical fusion expression vectors include pGEX (Pharmacia Biotech Inc; Smith, D.B. and Johnson, K.S. (1988) *Gene* 67:31-40), pMAL (New England Biolabs, Beverly, MA) and pRIT5 (Pharmacia, Piscataway, NJ) which fuse glutathione S-transferase (GST), maltose E binding protein, or protein A, respectively, to the target recombinant protein. In one embodiment, the coding sequence of the MP protein is cloned into a pGEX expression vector to create a vector encoding a fusion protein comprising, from

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the N-terminus to the C-terminus, GST-thrombin cleavage site-X protein. The fusion protein can be purified by affinity chromatography using glutathione-agarose resin. Recombinant MP protein unfused to GST can be recovered by cleavage of the fusion protein with thrombin.

Examples of suitable inducible non-fusion E. coli expression vectors include 5 pTrc (Amann et al., (1988) Gene 69:301-315) pLG338, pACYC184, pBR322, pUC18, pUC19, pKC30, pRep4, pHS1, pHS2, pPLc236, pMBL24, pLG200, pUR290, pIN-III113-B1, \(\lambda\)gt11, \(p\)BdCl, and \(p\)ET 11d (Studier et al., \(Gene\) Expression Technology: Methods in Enzymology 185, Academic Press, San Diego, California (1990) 60-89; and Pouwels et al., eds. (1985) Cloning Vectors. Elsevier: New York IBSN 0 444 904018). 10 Target gene expression from the pTrc vector relies on host RNA polymerase transcription from a hybrid trp-lac fusion promoter. Target gene expression from the pET 11d vector relies on transcription from a T7 gn10-lac fusion promoter mediated by a coexpressed viral RNA polymerase (T7 gn1). This viral polymerase is supplied by host strains BL21(DE3) or HMS174(DE3) from a resident λ prophage harboring a T7 15 gn1 gene under the transcriptional control of the lacUV 5 promoter. For transformation of other varieties of bacteria, appropriate vectors may be selected. For example, the plasmids pIJ101, pIJ364, pIJ702 and pIJ361 are known to be useful in transforming Streptomyces, while plasmids pUB110, pC194, or pBD214 are suited for transformation of Bacillus species. Several plasmids of use in the transfer of genetic information into 20 Corynebacterium include pHM1519, pBL1, pSA77, or pAJ667 (Pouwels et al., eds. (1985) Cloning Vectors. Elsevier: New York IBSN 0 444 904018).

One strategy to maximize recombinant protein expression is to express the protein in a host bacteria with an impaired capacity to proteolytically cleave the recombinant protein (Gottesman, S., *Gene Expression Technology: Methods in Enzymology* 185, Academic Press, San Diego, California (1990) 119-128). Another strategy is to alter the nucleic acid sequence of the nucleic acid to be inserted into an expression vector so that the individual codons for each amino acid are those preferentially utilized in the bacterium chosen for expression, such as *C. glutamicum* (Wada *et al.* (1992) *Nucleic Acids Res.* 20:2111-2118). Such alteration of nucleic acid sequences of the invention can be carried out by standard DNA synthesis techniques.

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In another embodiment, the MP protein expression vector is a yeast expression vector. Examples of vectors for expression in yeast *S. cerevisiae* include pYepSec1 (Baldari, *et al.*, (1987) *Embo J.* 6:229-234), , 2 μ, pAG-1, Yep6, Yep13, pEMBLYe23, pMFa (Kurjan and Herskowitz, (1982) *Cell* 30:933-943), pJRY88 (Schultz *et al.*, (1987) *Gene* 54:113-123), and pYES2 (Invitrogen Corporation, San Diego, CA). Vectors and methods for the construction of vectors appropriate for use in other fungi, such as the filamentous fungi, include those detailed in: van den Hondel, C.A.M.J.J. & Punt, P.J. (1991) "Gene transfer systems and vector development for filamentous fungi, in: Applied Molecular Genetics of Fungi, J.F. Peberdy, *et al.*, eds., p. 1-28, Cambridge University Press: Cambridge, and Pouwels *et al.*, eds. (1985) Cloning Vectors. Elsevier: New York (IBSN 0 444 904018).

Alternatively, the MP proteins of the invention can be expressed in insect cells using baculovirus expression vectors. Baculovirus vectors available for expression of proteins in cultured insect cells (e.g., Sf 9 cells) include the pAc series (Smith et al. (1983) Mol. Cell Biol. 3:2156-2165) and the pVL series (Lucklow and Summers (1989) Virology 170:31-39).

In another embodiment, the MP proteins of the invention may be expressed in unicellular plant cells (such as algae) or in plant cells from higher plants (e.g., the spermatophytes, such as crop plants). Examples of plant expression vectors include those detailed in: Becker, D., Kemper, E., Schell, J. and Masterson, R. (1992) "New plant binary vectors with selectable markers located proximal to the left border", *Plant Mol. Biol.* 20: 1195-1197; and Bevan, M.W. (1984) "Binary *Agrobacterium* vectors for plant transformation", *Nucl. Acid. Res.* 12: 8711-8721, and include pLGV23, pGHlac+, pBIN19, pAK2004, and pDH51 (Pouwels et al., eds. (1985) Cloning Vectors. Elsevier: New York IBSN 0 444 904018).

In yet another embodiment, a nucleic acid of the invention is expressed in mammalian cells using a mammalian expression vector. Examples of mammalian expression vectors include pCDM8 (Seed, B. (1987) *Nature* 329:840) and pMT2PC (Kaufman *et al.* (1987) *EMBO J.* 6:187-195). When used in mammalian cells, the expression vector's control functions are often provided by viral regulatory elements. For example, commonly used promoters are derived from polyoma, Adenovirus 2, cytomegalovirus and Simian Virus 40. For other suitable expression systems for both

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prokaryotic and eukaryotic cells see chapters 16 and 17 of Sambrook, J., Fritsh, E. F., and Maniatis, T. *Molecular Cloning: A Laboratory Manual. 2nd, ed., Cold Spring Harbor Laboratory,* Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989.

In another embodiment, the recombinant mammalian expression vector is 5 capable of directing expression of the nucleic acid preferentially in a particular cell type (e.g., tissue-specific regulatory elements are used to express the nucleic acid). Tissuespecific regulatory elements are known in the art. Non-limiting examples of suitable tissue-specific promoters include the albumin promoter (liver-specific; Pinkert et al. (1987) Genes Dev. 1:268-277), lymphoid-specific promoters (Calame and Eaton (1988) 10 Adv. Immunol. 43:235-275), in particular promoters of T cell receptors (Winoto and Baltimore (1989) EMBO J. 8:729-733) and immunoglobulins (Banerji et al. (1983) Cell 33:729-740; Queen and Baltimore (1983) Cell 33:741-748), neuron-specific promoters (e.g., the neurofilament promoter; Byrne and Ruddle (1989) PNAS 86:5473-5477), pancreas-specific promoters (Edlund et al. (1985) Science 230:912-916), and mammary 15 gland-specific promoters (e.g., milk whey promoter; U.S. Patent No. 4,873,316 and European Application Publication No. 264,166). Developmentally-regulated promoters are also encompassed, for example the murine hox promoters (Kessel and Gruss (1990) Science 249:374-379) and the α -fetoprotein promoter (Campes and Tilghman (1989) Genes Dev. 3:537-546). 20

The invention further provides a recombinant expression vector comprising a

DNA molecule of the invention cloned into the expression vector in an antisense
orientation. That is, the DNA molecule is operatively linked to a regulatory sequence in
a manner which allows for expression (by transcription of the DNA molecule) of an
RNA molecule which is antisense to MP mRNA. Regulatory sequences operatively
linked to a nucleic acid cloned in the antisense orientation can be chosen which direct
the continuous expression of the antisense RNA molecule in a variety of cell types, for
instance viral promoters and/or enhancers, or regulatory sequences can be chosen which
direct constitutive, tissue specific or cell type specific expression of antisense RNA.

The antisense expression vector can be in the form of a recombinant plasmid, phagemid
or attenuated virus in which antisense nucleic acids are produced under the control of a
high efficiency regulatory region, the activity of which can be determined by the cell

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type into which the vector is introduced. For a discussion of the regulation of gene expression using antisense genes see Weintraub, H. et al., Antisense RNA as a molecular tool for genetic analysis, Reviews - Trends in Genetics, Vol. 1(1) 1986.

Another aspect of the invention pertains to host cells into which a recombinant expression vector of the invention has been introduced. The terms "host cell" and "recombinant host cell" are used interchangeably herein. It is understood that such terms refer not only to the particular subject cell but to the progeny or potential progeny of such a cell. Because certain modifications may occur in succeeding generations due to either mutation or environmental influences, such progeny may not, in fact, be identical to the parent cell, but are still included within the scope of the term as used herein.

A host cell can be any prokaryotic or eukaryotic cell. For example, an MP protein can be expressed in bacterial cells such as *C. glutamicum*, insect cells, yeast or mammalian cells (such as Chinese hamster ovary cells (CHO) or COS cells). Other suitable host cells are known to those of ordinary skill in the art. Microorganisms related to *Corynebacterium glutamicum* which may be conveniently used as host cells for the nucleic acid and protein molecules of the invention are set forth in Table 3.

Vector DNA can be introduced into prokaryotic or eukaryotic cells via conventional transformation or transfection techniques. As used herein, the terms "transformation" and "transfection", "conjugation" and "transduction" are intended to refer to a variety of art-recognized techniques for introducing foreign nucleic acid (e.g., linear DNA or RNA (e.g., a linearized vector or a gene construct alone without a vector) or nucleic acid in the form of a vector (e.g., a plasmid, phage, phasmid, phagemid, transposon or other DNA) into a host cell, including calcium phosphate or calcium chloride co-precipitation, DEAE-dextran-mediated transfection, lipofection, natural competence, chemical-mediated transfer, or electroporation. Suitable methods for transforming or transfecting host cells can be found in Sambrook, et al. (Molecular Cloning: A Laboratory Manual. 2nd, ed., Cold Spring Harbor Laboratory, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989), and other laboratory manuals.

For stable transfection of mammalian cells, it is known that, depending upon the expression vector and transfection technique used, only a small fraction of cells may integrate the foreign DNA into their genome. In order to identify and select these

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integrants, a gene that encodes a selectable marker (e.g., resistance to antibiotics) is generally introduced into the host cells along with the gene of interest. Preferred selectable markers include those which confer resistance to drugs, such as G418, hygromycin and methotrexate. Nucleic acid encoding a selectable marker can be introduced into a host cell on the same vector as that encoding an MP protein or can be introduced on a separate vector. Cells stably transfected with the introduced nucleic acid can be identified by drug selection (e.g., cells that have incorporated the selectable marker gene will survive, while the other cells die).

To create a homologous recombinant microorganism, a vector is prepared which contains at least a portion of an MP gene into which a deletion, addition or substitution 10 has been introduced to thereby alter, e.g., functionally disrupt, the MP gene. Preferably, this MP gene is a Corynebacterium glutamicum MP gene, but it can be a homologue from a related bacterium or even from a mammalian, yeast, or insect source. In a preferred embodiment, the vector is designed such that, upon homologous recombination, the endogenous MP gene is functionally disrupted (i.e., no longer 15 encodes a functional protein; also referred to as a "knock out" vector). Alternatively, the vector can be designed such that, upon homologous recombination, the endogenous MP gene is mutated or otherwise altered but still encodes functional protein (e.g., the upstream regulatory region can be altered to thereby alter the expression of the endogenous MP protein). In the homologous recombination vector, the altered portion 20 of the MP gene is flanked at its 5' and 3' ends by additional nucleic acid of the MP gene to allow for homologous recombination to occur between the exogenous MP gene carried by the vector and an endogenous MP gene in a microorganism. The additional flanking MP nucleic acid is of sufficient length for successful homologous recombination with the endogenous gene. Typically, several kilobases of flanking DNA (both at the 5' and 3' ends) are included in the vector (see e.g., Thomas, K.R., and Capecchi, M.R. (1987) Cell 51: 503 for a description of homologous recombination vectors). The vector is introduced into a microorganism (e.g., by electroporation) and cells in which the introduced MP gene has homologously recombined with the endogenous MP gene are selected, using art-known techniques. 30

In another embodiment, recombinant microorganisms can be produced which contain selected systems which allow for regulated expression of the introduced gene.

For example, inclusion of an MP gene on a vector placing it under control of the lac operon permits expression of the MP gene only in the presence of IPTG. Such regulatory systems are well known in the art.

In another embodiment, an endogenous MP gene in a host cell is disrupted (e.g., by homologous recombination or other genetic means known in the art) such that expression of its protein product does not occur. In another embodiment, an endogenous or introduced MP gene in a host cell has been altered by one or more point mutations, deletions, or inversions, but still encodes a functional MP protein. In still another embodiment, one or more of the regulatory regions (e.g., a promoter, repressor, or inducer) of an MP gene in a microorganism has been altered (e.g., by deletion, truncation, inversion, or point mutation) such that the expression of the MP gene is modulated. One of ordinary skill in the art will appreciate that host cells containing more than one of the described MP gene and protein modifications may be readily produced using the methods of the invention, and are meant to be included in the present invention.

A host cell of the invention, such as a prokaryotic or eukaryotic host cell in culture, can be used to produce (*i.e.*, express) an MP protein. Accordingly, the invention further provides methods for producing MP proteins using the host cells of the invention. In one embodiment, the method comprises culturing the host cell of invention (into which a recombinant expression vector encoding an MP protein has been introduced, or into which genome has been introduced a gene encoding a wild-type or altered MP protein) in a suitable medium until MP protein is produced. In another embodiment, the method further comprises isolating MP proteins from the medium or the host cell.

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C. Isolated MP Proteins

Another aspect of the invention pertains to isolated MP proteins, and biologically active portions thereof. An "isolated" or "purified" protein or biologically active portion thereof is substantially free of cellular material when produced by recombinant DNA techniques, or chemical precursors or other chemicals when chemically synthesized. The language "substantially free of cellular material" includes preparations of MP protein in which the protein is separated from cellular components of the cells in which

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it is naturally or recombinantly produced. In one embodiment, the language "substantially free of cellular material" includes preparations of MP protein having less than about 30% (by dry weight) of non-MP protein (also referred to herein as a "contaminating protein"), more preferably less than about 20% of non-MP protein, still more preferably less than about 10% of non-MP protein, and most preferably less than about 5% non-MP protein. When the MP protein or biologically active portion thereof is recombinantly produced, it is also preferably substantially free of culture medium, i.e., culture medium represents less than about 20%, more preferably less than about 10%, and most preferably less than about 5% of the volume of the protein preparation. The language "substantially free of chemical precursors or other chemicals" includes preparations of MP protein in which the protein is separated from chemical precursors or other chemicals which are involved in the synthesis of the protein. In one embodiment, the language "substantially free of chemical precursors or other chemicals" includes preparations of MP protein having less than about 30% (by dry weight) of chemical precursors or non-MP chemicals, more preferably less than about 20% chemical precursors or non-MP chemicals, still more preferably less than about 10% chemical precursors or non-MP chemicals, and most preferably less than about 5% chemical precursors or non-MP chemicals. In preferred embodiments, isolated proteins or biologically active portions thereof lack contaminating proteins from the same organism from which the MP protein is derived. Typically, such proteins are produced by recombinant expression of, for example, a C. glutamicum MP protein in a microorganism such as C. glutamicum.

An isolated MP protein or a portion thereof of the invention can catalyze an enzymatic reaction in an amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathway, or has one or more of the activities set forth in Table 1. In preferred embodiments, the protein or portion thereof comprises an amino acid sequence which is sufficiently homologous to an amino acid sequence of the invention (e.g., a sequence of an even-numbered SEQ ID NO: of the Sequence Listing) such that the protein or portion thereof maintains the ability to catalyze an enzymatic reaction in an amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathway. The portion of the protein is preferably a biologically active portion as described herein. In another preferred embodiment, an MP protein of

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the invention has an amino acid sequence set forth as an even-numbered SEQ ID NO: of the Sequence Listing. In yet another preferred embodiment, the MP protein has an amino acid sequence which is encoded by a nucleotide sequence which hybridizes, e.g., hybridizes under stringent conditions, to a nucleotide sequence of the invention (e.g., a sequence of an odd-numbered SEQ ID NO: of the Sequence Listing). In still another preferred embodiment, the MP protein has an amino acid sequence which is encoded by a nucleotide sequence that is at least about 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, or 60%, preferably at least about 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, or 70%, more preferably at least about 71%, 72%, 73%, 74%, 75%, 76%, 77%, 78%, 79%, or 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, or 90%, or 91%, 92%, 93%, 94%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more homologous to one of the nucleic acid sequences of the invention, or a portion thereof. Ranges and identity values intermediate to the above-recited values, (e.g., 70-90% identical or 80-95% identical) are also intended to be encompassed by the present invention. For example, ranges of identity values using a combination of any of the above values recited as upper and/or lower limits are intended to be included. The preferred MP proteins of the present invention also preferably possess at least one of the MP activities described herein. For example, a preferred MP protein of the present invention includes an amino acid sequence encoded by a nucleotide sequence which hybridizes, e.g., hybridizes under stringent conditions, to a nucleotide sequence of the invention, and which can catalyze an enzymatic reaction in an amino acid, vitamin, cofactor, nutraceutical, nucleotide, nucleoside, or trehalose metabolic pathway, or which has one or more of the activities set forth in Table 1.

In other embodiments, the MP protein is substantially homologous to an amino acid sequence of the invention (*e.g.*, a sequence of an even-numbered SEQ ID NO: of the Sequence Listing) and retains the functional activity of the protein of one of the amino acid sequences of the invention yet differs in amino acid sequence due to natural variation or mutagenesis, as described in detail in subsection I above. Accordingly, in another embodiment, the MP protein is a protein which comprises an amino acid sequence which is at least about 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, or 60%, preferably at least about 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, or 70%, more preferably at least about 71%, 72%, 73%, 74%, 75%, 76%, 77%,

78%, 79%, or 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, or 90%, or 91%, 92%, 93%, 94%, and even more preferably at least about 95%, 96%, 97%, 98%, 99% or more homologous to an entire amino acid sequence of the invention and which has at least one of the MP activities described herein. Ranges and identity values intermediate to the above-recited values, (e.g., 70-90% identical or 80-95% identical) are also intended to be encompassed by the present invention. For example, ranges of identity values using a combination of any of the above values recited as upper and/or lower limits are intended to be included. In another embodiment, the invention pertains to a full length *C. glutamicum* protein which is substantially homologous to an entire amino acid sequence of the invention.

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Biologically active portions of an MP protein include peptides comprising amino acid sequences derived from the amino acid sequence of an MP protein, *e.g.*, an amino acid sequence of an even-numbered SEQ ID NO: of the Sequence Listing or the amino acid sequence of a protein homologous to an MP protein, which include fewer amino acids than a full length MP protein or the full length protein which is homologous to an MP protein, and exhibit at least one activity of an MP protein. Typically, biologically active portions (peptides, *e.g.*, peptides which are, for example, 5, 10, 15, 20, 30, 35, 36, 37, 38, 39, 40, 50, 100 or more amino acids in length) comprise a domain or motif with at least one activity of an MP protein. Moreover, other biologically active portions, in which other regions of the protein are deleted, can be prepared by recombinant techniques and evaluated for one or more of the activities described herein. Preferably, the biologically active portions of an MP protein include one or more selected domains/motifs or portions thereof having biological activity.

MP proteins are preferably produced by recombinant DNA techniques. For example, a nucleic acid molecule encoding the protein is cloned into an expression vector (as described above), the expression vector is introduced into a host cell (as described above) and the MP protein is expressed in the host cell. The MP protein can then be isolated from the cells by an appropriate purification scheme using standard protein purification techniques. Alternative to recombinant expression, an MP protein, polypeptide, or peptide can be synthesized chemically using standard peptide synthesis techniques. Moreover, native MP protein can be isolated from cells (e.g., endothelial

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cells), for example using an anti-MP antibody, which can be produced by standard techniques utilizing an MP protein or fragment thereof of this invention.

The invention also provides MP chimeric or fusion proteins. As used herein, an MP "chimeric protein" or "fusion protein" comprises an MP polypeptide operatively linked to a non-MP polypeptide. An "MP polypeptide" refers to a polypeptide having an amino acid sequence corresponding to MP, whereas a "non-MP polypeptide" refers to a polypeptide having an amino acid sequence corresponding to a protein which is not substantially homologous to the MP protein, e.g., a protein which is different from the MP protein and which is derived from the same or a different organism. Within the fusion protein, the term "operatively linked" is intended to indicate that the MP polypeptide and the non-MP polypeptide are fused in-frame to each other. The non-MP polypeptide can be fused to the N-terminus or C-terminus of the MP polypeptide. For example, in one embodiment the fusion protein is a GST-MP fusion protein in which the MP sequences are fused to the C-terminus of the GST sequences. Such fusion proteins can facilitate the purification of recombinant MP proteins. In another embodiment, the fusion protein is an MP protein containing a heterologous signal sequence at its Nterminus. In certain host cells (e.g., mammalian host cells), expression and/or secretion of an MP protein can be increased through use of a heterologous signal sequence.

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Preferably, an MP chimeric or fusion protein of the invention is produced by standard recombinant DNA techniques. For example, DNA fragments coding for the different polypeptide sequences are ligated together in-frame in accordance with conventional techniques, for example by employing blunt-ended or stagger-ended termini for ligation, restriction enzyme digestion to provide for appropriate termini, filling-in of cohesive ends as appropriate, alkaline phosphatase treatment to avoid undesirable joining, and enzymatic ligation. In another embodiment, the fusion gene can be synthesized by conventional techniques including automated DNA synthesizers. Alternatively, PCR amplification of gene fragments can be carried out using anchor primers which give rise to complementary overhangs between two consecutive gene fragments which can subsequently be annealed and reamplified to generate a chimeric gene sequence (see, for example, *Current Protocols in Molecular Biology*, eds. Ausubel *et al.* John Wiley & Sons: 1992). Moreover, many expression vectors are commercially available that already encode a fusion moiety (*e.g.*, a GST polypeptide). An MP-

encoding nucleic acid can be cloned into such an expression vector such that the fusion moiety is linked in-frame to the MP protein.

Homologues of the MP protein can be generated by mutagenesis, *e.g.*, discrete point mutation or truncation of the MP protein. As used herein, the term "homologue" refers to a variant form of the MP protein which acts as an agonist or antagonist of the activity of the MP protein. An agonist of the MP protein can retain substantially the same, or a subset, of the biological activities of the MP protein. An antagonist of the MP protein can inhibit one or more of the activities of the naturally occurring form of the MP protein, by, for example, competitively binding to a downstream or upstream member of the MP cascade which includes the MP protein. Thus, the *C. glutamicum* MP protein and homologues thereof of the present invention may modulate the activity of one or more metabolic pathways in which MP proteins play a role in this microorganism.

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In an alternative embodiment, homologues of the MP protein can be identified by screening combinatorial libraries of mutants, e.g., truncation mutants, of the MP 15 protein for MP protein agonist or antagonist activity. In one embodiment, a variegated library of MP variants is generated by combinatorial mutagenesis at the nucleic acid level and is encoded by a variegated gene library. A variegated library of MP variants can be produced by, for example, enzymatically ligating a mixture of synthetic 20 oligonucleotides into gene sequences such that a degenerate set of potential MP sequences is expressible as individual polypeptides, or alternatively, as a set of larger fusion proteins (e.g., for phage display) containing the set of MP sequences therein. There are a variety of methods which can be used to produce libraries of potential MP homologues from a degenerate oligonucleotide sequence. Chemical synthesis of a degenerate gene sequence can be performed in an automatic DNA synthesizer, and the 25 synthetic gene then ligated into an appropriate expression vector. Use of a degenerate set of genes allows for the provision, in one mixture, of all of the sequences encoding the desired set of potential MP sequences. Methods for synthesizing degenerate oligonucleotides are known in the art (see, e.g., Narang, S.A. (1983) Tetrahedron 39:3; 30 Itakura et al. (1984) Annu. Rev. Biochem. 53:323; Itakura et al. (1984) Science 198:1056; Ike et al. (1983) Nucleic Acid Res. 11:477.

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In addition, libraries of fragments of the MP protein coding can be used to generate a variegated population of MP fragments for screening and subsequent selection of homologues of an MP protein. In one embodiment, a library of coding sequence fragments can be generated by treating a double stranded PCR fragment of an MP coding sequence with a nuclease under conditions wherein nicking occurs only about once per molecule, denaturing the double stranded DNA, renaturing the DNA to form double stranded DNA which can include sense/antisense pairs from different nicked products, removing single stranded portions from reformed duplexes by treatment with S1 nuclease, and ligating the resulting fragment library into an expression vector. By this method, an expression library can be derived which encodes N-terminal, C-terminal and internal fragments of various sizes of the MP protein.

Several techniques are known in the art for screening gene products of combinatorial libraries made by point mutations or truncation, and for screening cDNA libraries for gene products having a selected property. Such techniques are adaptable for rapid screening of the gene libraries generated by the combinatorial mutagenesis of MP homologues. The most widely used techniques, which are amenable to high through-put analysis, for screening large gene libraries typically include cloning the gene library into replicable expression vectors, transforming appropriate cells with the resulting library of vectors, and expressing the combinatorial genes under conditions in which detection of a desired activity facilitates isolation of the vector encoding the gene whose product was detected. Recursive ensemble mutagenesis (REM), a new technique which enhances the frequency of functional mutants in the libraries, can be used in combination with the screening assays to identify MP homologues (Arkin and Yourvan (1992) PINAS 89:7811-7815; Delgrave et al. (1993) Protein Engineering 6(3):327-331).

In another embodiment, cell based assays can be exploited to analyze a variegated MP library, using methods well known in the art.

D. Uses and Methods of the Invention

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The nucleic acid molecules, proteins, protein homologues, fusion proteins, primers, vectors, and host cells described herein can be used in one or more of the following methods: identification of *C. glutamicum* and related organisms; mapping of genomes of organisms related to *C. glutamicum*; identification and localization of *C.*

glutamicum sequences of interest; evolutionary studies; determination of MP protein regions required for function; modulation of an MP protein activity; modulation of the activity of an MP pathway; and modulation of cellular production of a desired compound, such as a fine chemical.

5 The MP nucleic acid molecules of the invention have a variety of uses. First, they may be used to identify an organism as being Corynebacterium glutamicum or a close relative thereof. Also, they may be used to identify the presence of C. glutamicum or a relative thereof in a mixed population of microorganisms. The invention provides the nucleic acid sequences of a number of C. glutamicum genes; by probing the 10 extracted genomic DNA of a culture of a unique or mixed population of microorganisms under stringent conditions with a probe spanning a region of a C. glutamicum gene which is unique to this organism, one can ascertain whether this organism is present. Although Corynebacterium glutamicum itself is not pathogenic to humans, it is related to species which are human pathogens, such as Corynebacterium diphtheriae. 15 Corynebacterium diphtheriae is the causative agent of diphtheria, a rapidly developing, acute, febrile infection which involves both local and systemic pathology. In this disease, a local lesion develops in the upper respiratory tract and involves necrotic injury to epithelial cells; the bacilli secrete toxin which is disseminated through this lesion to distal susceptible tissues of the body. Degenerative changes brought about by the 20 inhibition of protein synthesis in these tissues, which include heart, muscle, peripheral nerves, adrenals, kidneys, liver and spleen, result in the systemic pathology of the disease. Diphtheria continues to have high incidence in many parts of the world, including Africa, Asia, Eastern Europe and the independent states of the former Soviet Union. An ongoing epidemic of diphtheria in the latter two regions has resulted in at 25 least 5,000 deaths since 1990.

In one embodiment, the invention provides a method of identifying the presence or activity of *Cornyebacterium diphtheriae* in a subject. This method includes detection of one or more of the nucleic acid or amino acid sequences of the invention (e.g., the sequences set forth as odd-numbered or even-numbered SEQ ID NOs, respectively, in the Sequence Listing) in a subject, thereby detecting the presence or activity of *Corynebacterium diphtheriae* in the subject. *C. glutamicum* and *C. diphtheriae* are related bacteria, and many of the nucleic acid and protein molecules in *C. glutamicum*

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are homologous to *C. diphtheriae* nucleic acid and protein molecules, and can therefore be used to detect *C. diphtheriae* in a subject.

The nucleic acid and protein molecules of the invention may also serve as markers for specific regions of the genome. This has utility not only in the mapping of the genome, but also for functional studies of *C. glutamicum* proteins. For example, to identify the region of the genome to which a particular *C. glutamicum* DNA-binding protein binds, the *C. glutamicum* genome could be digested, and the fragments incubated with the DNA-binding protein. Those which bind the protein may be additionally probed with the nucleic acid molecules of the invention, preferably with readily detectable labels; binding of such a nucleic acid molecule to the genome fragment enables the localization of the fragment to the genome map of *C. glutamicum*, and, when performed multiple times with different enzymes, facilitates a rapid determination of the nucleic acid sequence to which the protein binds. Further, the nucleic acid molecules of the invention may be sufficiently homologous to the sequences of related species such that these nucleic acid molecules may serve as markers for the construction of a genomic map in related bacteria, such as *Brevibacterium lactofermentum*.

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The MP nucleic acid molecules of the invention are also useful for evolutionary and protein structural studies. The metabolic processes in which the molecules of the invention participate are utilized by a wide variety of prokaryotic and eukaryotic cells; by comparing the sequences of the nucleic acid molecules of the present invention to those encoding similar enzymes from other organisms, the evolutionary relatedness of the organisms can be assessed. Similarly, such a comparison permits an assessment of which regions of the sequence are conserved and which are not, which may aid in determining those regions of the protein which are essential for the functioning of the enzyme. This type of determination is of value for protein engineering studies and may give an indication of what the protein can tolerate in terms of mutagenesis without losing function.

Manipulation of the MP nucleic acid molecules of the invention may result in the production of MP proteins having functional differences from the wild-type MP proteins. These proteins may be improved in efficiency or activity, may be present in greater numbers in the cell than is usual, or may be decreased in efficiency or activity.

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The invention also provides methods for screening molecules which modulate the activity of an MP protein, either by interacting with the protein itself or a substrate or binding partner of the MP protein, or by modulating the transcription or translation of an MP nucleic acid molecule of the invention. In such methods, a microorganism expressing one or more MP proteins of the invention is contacted with one or more test compounds, and the effect of each test compound on the activity or level of expression of the MP protein is assessed.

When the desired fine chemical to be isolated from large-scale fermentative culture of C. glutamicum is an amino acid, a vitamin, a cofactor, a nutraceutical, a nucleotide, a nucleoside, or trehalose, modulation of the activity or efficiency of activity of one or more of the proteins of the invention by recombinant genetic mechanisms may directly impact the production of one of these fine chemicals. For example, in the case of an enzyme in a biosynthetic pathway for a desired amino acid, improvement in efficiency or activity of the enzyme (including the presence of multiple copies of the gene) should lead to an increased production or efficiency of production of that desired amino acid. In the case of an enzyme in a biosynthetic pathway for an amino acid whose synthesis is in competition with the synthesis of a desired amino acid, any decrease in the efficiency or activity of this enzyme (including deletion of the gene) should result in an increase in production or efficiency of production of the desired amino acid, due to decreased competition for intermediate compounds and/or energy. In the case of an enzyme in a degradation pathway for a desired amino acid, any decrease in efficiency or activity of the enzyme should result in a greater yield or efficiency of production of the desired product due to a decrease in its degradation. Lastly, mutagenesis of an enzyme involved in the biosynthesis of a desired amino acid such that this enzyme is no longer is capable of feedback inhibition should result in increased yields or efficiency of production of the desired amino acid. The same should apply to the biosynthetic and degradative enzymes of the invention involved in the metabolism of vitamins, cofactors, nutraceuticals, nucleotides, nucleosides and trehalose.

Similarly, when the desired fine chemical is not one of the aforementioned compounds, the modulation of activity of one of the proteins of the invention may still impact the yield and/or efficiency of production of the compound from large-scale culture of *C. glutamicum*. The metabolic pathways of any organism are closely

interconnected; the intermediate used by one pathway is often supplied by a different pathway. Enzyme expression and function may be regulated based on the cellular levels of a compound from a different metabolic process, and the cellular levels of molecules necessary for basic growth, such as amino acids and nucleotides, may critically affect the viability of the microorganism in large-scale culture. Thus, modulation of an amino acid biosynthesis enzyme, for example, such that it is no longer responsive to feedback inhibition or such that it is improved in efficiency or turnover may result in increased cellular levels of one or more amino acids. In turn, this increased pool of amino acids provides not only an increased supply of molecules necessary for protein synthesis, but also of molecules which are utilized as intermediates and precursors in a number of other biosynthetic pathways. If a particular amino acid had been limiting in the cell, its increased production might increase the ability of the cell to perform numerous other metabolic reactions, as well as enabling the cell to more efficiently produce proteins of all kinds, possibly increasing the overall growth rate or survival ability of the cell in large scale culture. Increased viability improves the number of cells capable of producing the desired fine chemical in fermentative culture, thereby increasing the yield of this compound. Similar processes are possible by the modulation of activity of a degradative enzyme of the invention such that the enzyme no longer catalyzes, or catalyzes less efficiently, the degradation of a cellular compound which is important for the biosynthesis of a desired compound, or which will enable the cell to grow and reproduce more efficiently in large-scale culture. It should be emphasized that optimizing the degradative activity or decreasing the biosynthetic activity of certain molecules of the invention may also have a beneficial effect on the production of certain fine chemicals from C. glutamicum. For example, by decreasing the efficiency of activity of a biosynthetic enzyme in a pathway which competes with the biosynthetic pathway of a desired compound for one or more intermediates, more of those intermediates should be available for conversion to the desired product. A similar situation may call for the improvement of degradative ability or efficiency of one or more proteins of the invention.

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This aforementioned list of mutagenesis strategies for MP proteins to result in increased yields of a desired compound is not meant to be limiting; variations on these mutagenesis strategies will be readily apparent to one of ordinary skill in the art. By

these mechanisms, the nucleic acid and protein molecules of the invention may be utilized to generate *C. glutamicum* or related strains of bacteria expressing mutated MP nucleic acid and protein molecules such that the yield, production, and/or efficiency of production of a desired compound is improved. This desired compound may be any natural product of *C. glutamicum*, which includes the final products of biosynthesis pathways and intermediates of naturally-occurring metabolic pathways, as well as molecules which do not naturally occur in the metabolism of *C. glutamicum*, but which are produced by a *C. glutamicum* strain of the invention.

This invention is further illustrated by the following examples which should not be construed as limiting. The contents of all references, patent applications, patents, published patent applications, Tables, and the sequence listing cited throughout this application are hereby incorporated by reference.

TABLE 1: Included Genes

Lysine biosynthesis

NT Start NT Stop Function 2793 3617 DIAMINOPIMELATE EPIMERASE (EC 5.1.1.7) ACETYLORNITHINE AMINOTRANSFERASE (EC 2.6.1.11) ACETYLORNITHINE AMINOTRANSFERASE (EC 2.6.1.11) MEMBRANE SPANNING PROTEIN INVOLVED IN LYSINE METABOLISM CYTOSOLIC PROTEIN INVOLVED IN METABOLISM CYTOSOLIC PROTEIN INVOLVED IN METABOLISM CYTOSOLIC PROTEIN INVOLVED IN LYSINE METABOLISM	NT Start NT Stop Function 37078 38532 ALPHA,ALPHA-TREHALOSE-PHOSPHATE SYNTHASE (UDP-FORMING) 56 KD SUBUNIT (EC 2.4.1.15) 1486 2931 ALPHA,ALPHA,TREHALOSE-PHOSPHATE SYNTHASE (UDP-FORMING) 56 KD	SUBUNIT (EC 2 4 1.15) 3 758 trehalose synthase (EC 2.4.1) 1005 4 trehalose synthase (EC 2.4.1)	NT Start NT Stop Function 4758 3496 ASPARTOKINASE ALPHA AND BETA SUBUNITS (EC 2.7.2.4) 3469 2438 ASPARTATE-SEMIALDEHYDE DEHYDROGENASE (EC 1.2.1.11) 543 4 2,3.4.5-TETRAHYDROPYRIDINE-2-CARBOXYLATE N-SUCCINYLTRANSFERASE 2063 3169 SUCCINYL-DIAMINOPIMELATE DESUCCINYLASE (EC 3.5.1.18) 3458 4393 DIHYDRODIPICOLINATE SYNTHASE (EC 4.2.1.52) 1694 2443 probable 2,3-dihydrodipicolinate N-C6-lyase (cyclizing) (EC 4.3.3) - Corynebacterium glutamicum 2,3.4.5-TETRAHYDROPYRIDINE-2-CARBOXYLATE N-SUCCINYLTRANSFERASE (EC 2.3.1.117) (EC 2.3.1.117) 31980 30961 MESO-DIAMINOPIMELATE D-DEHYDROGENASE MESO-DIAMINOPIMELATE D-DEHYDROGENASE
iart I	벌		±!
Contig. GR006 3 GR002 7	Contig. VV013	GR00241 GR00243	Contig GR00137 GR00642 GR00613 GR000236 GR00236 GR00236 GR00236 GR00236 GR00236
RXA02229 RXS02970 F RXA01009 RXC02390 RXC01796 RXC01207 RXC00657 RXC00655	Identification Code RXN00351 F RXA00351	XXA00873 XXA00891	ldentification Code RXA00534 RXA00533 RXA02843 RXA02022 RXA00044 RXA00863 RXA00864 RXA00864 RXA00855 F RXA00355
Amino Acid SEQ ID NO 2 4 6 6 6 12 12	Amino Acid SEO ID NO 18	22 E5 23 24 E Lysine biosynthesis	Amino Acid SEQ ID NO 28 28 30 33 34 40 40
Nucleic Acid SEQ ID NO 1 3 5 5 7 7 11	Trehalose Nucleic Acid SEQ ID NO 17	21 23 Lysine bi	Nucleic Acid SEQ ID NO 25 27 29 31 33 35 37 41

Table 1 (continued)	ode Contig. NT Start NT Stop Function		GR00274 3 1379 DIAMINOPIMELATE DECARBOXYLASE (EC 4.1.1.20)	GR00752 5237 7234 DIAMINOPIMELATE DECARBOXYLASE (EC 4.1.1.20)	GR00408 4249 3380 LYSINE EXPORT REGULATOR PROTEIN	GR00036 5443 6945 L-LYSINE TRANSPORT PROTEIN	GR00408 4320 5018 LYSINE EXPORTER PROTEIN	GR00236 2647 3549 DIHYDRODIPICOLINATE SYNTHASE (EC 4.2.1.52)	2,3,4,5-TETRAHYDROPYRIDINE-2-CARBOXYLATE N-SUCCINYLTRANSFERASE	(EC 2.3.1.117)	ACETYLORNITHINE AMINOTRANSFERASE (EC 2 6.1.11)	ABC TRANSPORTER ATP-BINDING PROTEIN INVOLVED IN LYSINE	METABOLISM	PROTEIN INVOLVED IN LYSINE METABOLISM	ZN-DEPENDENT HYDROLASE INVOLVED IN LYSINE METABOLISM	ABC TRANSPORTER ATP-BINDING PROTEIN INVOLVED IN LYSINE	METABOLISM	PROTEIN (NVOLVED IN LYSINE METABOLISM
	Identification Code Contig.		RXA00972	RXA02653	RXA01393	RXA00241	RXA01394	RXA00865	RXS02021		RXS02157	RXC00733		RXC00861	RXC00866	RXC02095		RXC03185
	Amino Acid	SEQ ID NO	46	48	20	52	54	26	58		90	62		64	99	99		70
	Nucleic Acid	SEQ ID NO	45	47	49	51	53	55	22		59	61		63	65	29		69

Glutamate and glutamine metabolism

Function	GLUTAMATE SYNTHASE INADHI PRECURSOR (EC 1.4.1.14)	GLUTAMATE SYNTHASE (NADPH) LARGE CHAIN PRECURSOR (EC 1.4.1.13)	GLUTAMATE SYNTHASE (NADPH) LARGE CHAIN PRECURSOR (EC 1,4,1,13)	GLUTAMATE SYNTHASE (NADPH) LARGE CHAIN PRECURSOR (EC 1.4.1.13)	GLUTAMATE SYNTHASE (NADPH) SMALL CHAIN (EC 1.4.1.13)	GLUTAMATE SYNTHASE [NADPH] SMALL CHAIN (EC 1.4.1.13)	GLUTAMATE SYNTHASE (NADPH) SMALL CHAIN (EC 1.4.1.13)	GLUTAMATE SYNTHASE (NADPH) SMALL CHAIN (EC 1.4.1.13)	NADP-SPECIFIC GLUTAMATE DEHYDROGENASE (EC 1.4.1.4)	GLUTAMINE SYNTHETASE (EC 6.3.1.2)	GLUTAMINE SYNTHETASE (EC 6.3.1.2)	GLUTAMATE-AMMONIA-LIGASE ADENYLYLTRANSFERASE (EC 2.7.7.42)	GLUTAMINASE (EC 3.5.1.2)	GLUTAMINASE (EC 3.5.1.2)	GLUTAMINE-BINDING PROTEIN PRECURSOR	GLUTAMINE-BINDING PERIPLASMIC PROTEIN PRECURSOR			
NT Stop	14273	8912	4	964	4122	3419	7368	283	15233	4	605	2599	5192	17750	8396	862	862	1581	1525
NT Start	9744	7107	1296	1806	2752	2757	7916	2	14607	630	961	1259	3855	19180	5262	2	2	2612	614
Contig	W0196	GR00001	GR00074	GR00075	VV0154	GR00012	W0181	GR00031	W0196	GR00075	GR00075	GR00628	GR00057	GR00057	GR00057	VV0332	GR10017	GR00043	GR00193
Identification Code	RXN00367	F RXA00007	F RXA00364	F RXA00367	RXN00076	F RXA00075	RXN00198	F RXA00198	RXN00365	F RXA00365	RXA00366	RXA02072	RXA00323	RXA00335	RXA00324	RXN03176	F RXA02879	RXA00278	RXA00727
Amino Acid SEQ ID NO	72	74	76	78	80	82	84	98	88	06	35	94	96	86	100	102	104	106	108
Nucleic Acid SEQ ID NO	71	73	75	77	79	81	83	85	87	89	91	93	96	26	66	101	103	105	107

Table 1 (continued) aragine metabolism	NT Start NT Stop Function 6739 4901 ASPARAGINE SYNTHETASE (GLUTAMINE-HYDROLYZING) (EC 6.3.5.4) 26974 25814 ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1) 10288 25814 ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1) 10289 3182 ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1) 854 1138 ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1) 1585 275 ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1) 2669 1695 L-ASPARAGINASE (EC 3.5.1.1) 680 6 ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1) 4701 5783 ALANINE RACEMASE (EC 5.1.1.1) 20972 19944 ALANINE RACEMASE, BIOSYNTHETIC (EC 5.1.1.1)	NT Start NT Stop Function 8581 7826 BETA-UREIDOPROPIONASE (EC 3.5.1.6) METHYLMALONATE-SEMIALDEHYDE DEHYDROGENASE (ACYLATING) (EC 1.2.1.27) ASPARTATE 1-DECARBOXYLASE PRECURSOR (EC 4.1.1.11)	NT Stap Function 1113 2042 L-SERINE DEHYDRATASE (EC 4.2.1.13) 481 1827 L-SERINE DEHYDRATASE (EC 4.2.1.13) 7343 6042 SERINE HYDROXYMETHYLTRANSFERASE (EC 2.1.2.1) 10253 9876 SARCOSINE OXIDASE (EC 1.5.3.1) 33454 33813 SARCOSINE OXIDASE (EC 1.5.3.1) 33454 12581 PHOSPHOSERINE PHOSPHATASE (EC 2.1.52) PHOSPHOSERINE PHOSPHATASE (EC 3.1.3.3) PHOSPHOSERINE PHOSPHATASE (EC 3.1.3.3) 5082 4648 PHOSPHOSERINE PHOSPHATASE (EC 3.1.3.3) 5084 PHOSPHOSERINE PHOSPHATASE (EC 3.1.3.3) 5520 PHOSPHOSERINE PHOSPHATASE (EC 3.1.3.3) 1543 SARCOSINE OXIDASE (EC 1.5.3.1) 15423 SARCOSINE OXIDASE (EC 1.5.3.1) 15423 SARCOSINE OXIDASE (EC 1.5.3.1) 15429 D-3-PHOSPHOGLYCERATE DEHYDROGENASE (EC 1.1.1.95)
ragine m	Contig. GR00639 VV0100 GR00018 VV0135 GR00163 GR00729 GR00729 GR00729 VV0138 VV0135	Contig. GR00726	Contig. GR00435 GR00525 GR005156 GR00515 VV0202 GR00654 GR00766 GR007766 GR007766 GR007766
Alanine and Aspartate and Aspai	Identification Code RXA02139 RXN00116 F RXA00116 F RXA00618 F RXA00618 F RXA00627 RXA02550 RXA02250 RXA02193 RXA02432 RXN03003 RXN03003 RXN00636	beta-Alanine metabolism Nucleic Acid SEQ ID NO 134 RXA02536 135 136 RXS02299 137 138 RXS02299 Glycine and serine metabolism	Identification Code RXA01561 RXA01850 RXA00580 RXA00263 F RXA02263 F RXA02263 F RXA02176 RXN02758 F RXA02759 F RXA02759 F RXA02759 F RXA02759 RXA02759 RXA027501 RXN03105 RXS01130 RXS01130
ınd Asparı	Amino Acid SEQ ID NO 110 112 118 118 120 122 124 126 130 132	beta-Alanine metabolism Nucleic Acid SEQ ID NO 133 134 RX 8008 137 138 RX 8008 Glycine and serine metak	Amino Acid SEQ ID NO 140 144 144 146 150 152 154 156 160 160 166
Alanine a	Nucleic Acid SEQ ID NO 109 111 113 115 117 119 121 123 125 127 131	Nucleic Acid SEQ ID NO 133 135 137 Glycine a	Nucleic Acid SEQ ID NO 139 141 143 145 149 151 153 155 161 163 167

Threonine metabolism

Table 1 (continued)

Function	HOMOSERINE DEHYDROGENASE (EC 1.1.1.3)	HOMOSERINE DEHYDROGENASE (EC 1.1.1.3)	HOMOSERINE KINASE (EC 2.7.1.39)	THREONINE SYNTHASE (EC 4.2.99.2)	HOMOSERINE O-ACETYLTRANSFERASE	HOMOSERINE O-ACETYLTRANSFERASE (EC 2.3.1.11)	CYTOSOLIC PROTEIN INVOLVED IN METABOLISM OF LYSINE AND	THREONINE	MEMBRANE ASSOCIATED PROTEIN INVOLVED IN THREONINE METABOLISM
art NT Stop	13387	3015	1087	14410	68911	1832			
NT Start	12053	2623	161	12968	70041	723			
Contig.	VV0149	GR00274	GR00273	GR00057	00000	GR00088			
Identification Code	_	F RXA00974	RXA00970	RXA00330	RXN00403	F RXA00403	RXC01207		RXC00152
Amino Acid SEQ ID NO		172	174	176	178	180	182		184
Nucleic Acid SEQ ID NO	169	171	173	175	177	179	181		183

Metabolism of methionine and S-adenosyl methionine

		se) (INE	SINE	NE	3E	SE.	χ.			
Function HOMOSERINE O-ACETYLTRANSFERASE (EC 2 3.1.31) HOMOSERINE O-ACETYLTRANSFERASE HOMOSERINE O-ACETYLTRANSFERASE (EC 2.3.1.11) CYSTATHIONINE GAMMA-SYNTHASE (EC 4 2 99 9)	CYSTATHIONINE GAMMA-SYNTHASE (EC 4.2 99 9) CYSTATHIONINE GAMMA-SYNTHASE (EC 4.2 99 9) CYSTATHIONINE GAMMA-SYNTHASE (EC 4.2 99 9)	OTSTATRIONINE GAMMIA-STNTHASE (EC. 4.2.99.9) 5-methyltetrahydrofoate-homocysteine methyltransferase (methionine synthetase) OACETYLHOMOSERINE SULFHYDRYLASE (EC. 4.2.99.10) / O-ACETYLSERINE SUIFHYDRYLASE (FC. 4.2.99.10)	O-ACETYLHOMOSERINE SULFHYDRYLASE (EC 4.2.99.10) / O-ACETYLSERINE SULFHYDRYLASE (EC 4.2.99.8)	O-ACETYLHOMOSERINE SULFHYDRYLASE (EC 4.2.99.10) / O-ACETYLSERINE SULFHYDRYLASE (EC 4.2.99.8)	5-METHYLTETRAHYDROFOLATEHOMOCYSTEINE METHYLTRANSFERASE (EC 2.1.1.13)	5-METHYLTETRAHYDROFOLATEHOMOCYSTEINE METHYLTRANSFERASE (EC 2.1.1.13)	5-METHYLTETRAHYDROFOLATEHOMOCYSTEINE METHYLTRANSFERASE (EC 2.1.1.13)	S-ADENOSYLMETHIONINE:2-DEMETHYLMENAQUINONE METHYLTRANSFERASE (EC 2.1.⊹.)	S-ADENOSYLMETHIONINE:2-DEMETHYLMENAQUINONE METHYLTRANSFERASE (EC 2 1 →)	ADENOSYLHOMOCYSTEINASE (EC 3.3.1.1) ADENOSYLHOMOCYSTEINASE (EC 3.3.1.1)
Function HOMOSERIN HOMOSERIN HOMOSERIN CYSTATHION	CYSTATHION CYSTATHION CYSTATHION	5-methyltetral O-ACETYLHC	O-ACETYLHO SULFHYDRYI	O-ACETYLHO SULFHYDRYI	5-METHYLTE (EC 2 1 1.13)	5-METHYLTE (EC 2.1.1.13)	5-METHYLTE (EC 2.1.1.13)	S-ADENOSYL METHYLTRAI	S-ADENOSYL METHYLTRAI	ADENOSYLH ADENOSYLH
NT Stop 4313 68911 1832	2039	2521 15297 70188	576	3801	4025	11726	9	1741	645	5045 7624
NT Start 5359 70041 723	2404 3085	16286 70787	-	3289	4552	9228	2483	2238	1142	3612 7728
Contig. GR00017 VV0086 GR00088	GR00038 GR00726	GR00032 VV0086	GR00088	GR00089	GR00645	VV0302	GR00646	VV0042	GR10044	VV0124 GR00020
Identification Code RXA00115 RXN00403 F RXA00403	F RXA00254 RXA02532 RXS03159 E BXA03758	RXA00216 RXN00402	F RXA00402	RXA00405	RXA02197	RXN02198	F RXA02198	RXN03074	F RXA02906	RXN00132 F RXA00132
Amino Acid SEQ ID NO 186 188 190	194 196 198	202 204 204	206	208	210	212	214	216	218	220 222
Nucleic Acid SEQ ID NO 185 187 191	193 195 197	201 203	205	207	209	211	213	215	217	219 221

tinued) Function	ADENOSYLHOMOCYSTEINASE (EC 3.3.1.1) 5-METHYLTETRAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTEINE METHYLTDANGERDAGE (EC. 3.4.4.1)	ME INTLINANSFERASE (EC. Z.T.T.14) S-METHYLTETRAHYDRODFEROYLFRIGLUTAMATEHOMOCYSTEINE METHYLTETRAHYDRODE (C. Z.T.T.14)	METHYLLERANSTERASE (EC. 2.1.1.14) 5-METHYLLETRAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTEINE METHYLTERANSTERASE (CC. 3.4.4.4)	METHYLLETRAHYDROPTEROOF (C. Z.1.).14) 5-METHYLTETRAHYDROPTEROOFTRIGLUTAMATEHOMOCYSTEINE METHYLTDANGEDAGE (C. Z. 2. 4. 4.4.)	S-METHYLLERANSPERASE (EC. 2.1.1.14) 5-METHYLLERAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTEINE	ME INTLIMANSERASE (EC 27.1.14) 5-METHYLTERHYDDOPTEROYLTRIGLUTAMATEHOMOCYSTEINE	ME INTLIKANSFEKASE (EU Z.1.3.14) PROTEIN INVED IN METABOLISM OF S-ADENOSYLMETHIONINE, PURINES	AND PANTOTHENATE EXPORTED PROTEIN INVOLVED IN METABOLISM OF PYRIDIMES AND ADENOSYLHOMOCYSTEINE		Function	S-ADENOSYLMETHIONINE SYNTHETASE (EC 2.5.1.6)		Function	SERINE ACETYLTRANSFERASE (EC 2.3.1.30)	CYSTEINE SYNTHASE (EC 4.2.99.8) O-ACETYLHOMOSERINE SULFHYDRYLASE (EC 4.2.99.10) / O-ACETYLSERINE	SULFHYDRYLASE (EC 4 2 99 8) O-ACETYLLMOMOSERINE SULFHYDRYLASE (EC 4 2 99 10) / O-ACETYLSERINE	SULFITURY LASE (EU 4.2.39.6) SULFITURY LASE (EU 4.2.39.10) / O-ACETYLSERINE SULFITURY ASE (EU 4.2.39.10) / O-ACETYLSERINE	SOLFHIUN LASE (EC. 4.2.39.0) METAPOLISM METAPOLISM	METABOLISM ABC TRANSPORTER ATP-BINDING PROTEIN INVOLVED IN CYSTEINE METABOLISM	
Table 1 (continued)	3634	5295	5731	_ u) =	4730	15447	e u •			NT Stop	8380		NT Stop	2234	m	576	,, ,			
Ta NT Start	2339	3496	5252		5254	14764			sis	NT Start	7160		NT Start	1689	550 70787	-				
Contig	GR00398	GR00629	GR00629		GR00751	GR00752			osynthe	Contig	GR00654		Contig	GR00206	GR00206 VV0086	GR00088				
Identification Code	F RXA01371 RXN02085	F RXA02085	F RXA02086	RXN02648	F RXA02648	F RXA02658	RXC02238	RXC00128	S-adenosyl methionine (SAM) Biosynthesis	Identification Code	RXA02240	E	Identification Code	RXA00780	RXA00779 RXN00402	F RXA00402	RXS00405	RXC00164	RXC01191	
Amino Acid	224 226	228	230	232	234	236	238	240	syl methio	Amino Acid	242	Cysteine metabolism	Amino Acid	244 244	246 248	250	252	254	256	
Ϋ́	0 0 0								<u>~</u>				Nucleic Acid							

Valine, leucine and isoleucine

Table 1 (continued)

Function	THREONINE DEHYDRATASE BIOSYNTHETIC (EC 4 2.1.16)	BRANCHED CHAIN AMINO ACID AMINOTRANSFERASE (EC 2.6.1.42)	BRANCHED-CHAIN AMINO ACID AMINOTRANSFERASE (EC 2.6.1.42)	BRANCHED-CHAIN AMINO ACID AMINOTRANSFERASE (EC 2.6.1.42)	3-ISOPROPYLMALATE DEHYDRATASE LARGE SUBUNIT (EC 4 2.1.33)	3-ISOPROPYLMALATE DEHYDRATASE LARGE SUBUNIT (EC 4.2.1.33)	3-ISOPROPYLMALATE DEHYDROGENASE (EC 1.1.1.85)	3-ISOPROPYLMALATE DEHYDROGENASE (EC 1-1-185)	2-ISOPROPYLMALATE SYNTHASE (EC 4 1 3 12)	2-ISOPROPYLMALATE SYNTHASE (EC 4.1.3.1)	3-ISOPROPYLMALATE DEHYDRATASE SMALL SUBUNIT (EC 4.2.1.33)	3-METHYL-2-OXOBUTANOATE HYDROXYMETHYLTRANSFERASE (EC 2.1.2.11)	/ DECARBOXYLASE (EC 4.1.144)	3-METHYL-2-OXOBUTANOATE HYDROXYMETHYLTRANSFERASE (EC 2.1.2.11)	4"-MYCAROSYL ISOVALERYL-COA TRANSFERASE (EC 2)	KETOL-ACID REDUCTOISOMERASE (EC 1.1.186)	KETOL-ACID REDUCTOISOMERASE (EC 1.1.1.86)
NT Stop	2588	4249	196	196	7513	1602	3472	1651	7498	7360	7121	48402		1960	14643		1530
NT Start	3856	5091	1296	1248	9171	-	4491	1349	6128	6128	7711	47590		2766	15584		1075
Contig.	GR00751	GR00204	VV0246	GR00473	VV0143	GR00294	VV0157	GR00315	VV0219	GR00137	VV0143	VV0127		GR00555	VV0122		GR00321
Identification Code	RXA02646	RXA00766	RXN01690	F RXA01690	RXN01026	F RXA01026	RXN01127	F RXA01132	RXN00536	F RXA00536	RXN02965	RXN01929		F RXA01929	RXN01420	RXS01145	F RXA01145
Amino Acid SEQ ID NO	258	260	262	264	266	268	270	272	274	276	278	280		282	284	286	288
Nucleic Acid	257											279					

Arginine and proline metabolism

Enzymes of proline biosynthesis:

unction	GLUTAMATE 5-KINASE (EC 2.7.2.11)	GAMMA-GLUTAMYL PHOSPHATE REDUCTASE (GPR) (EC. 1.2.1.41)	SAMMA-GLUTAMYL PHOSPHATE REDUCTASE (GPR) (EC 1.2.1.41)	PYRROLINE-5-CARBOXYLATE REDUCTASE (EC 1.5.1.2)	ACETYLORNITHINE AMINOTRANSFERASE (EC 2.6.1.11)	DRNITHINE CYCLODEAMINASE (EC 4.3.1.12)	ACETYLORNITHINE AMINOTRANSFERASE (EC 2.6.1.11)	ACETYLORNITHINE AMINOTRANSFERASE (EC 2 6 1 11)
NT Start NT Stop Function	223	3607 16	1894	12692				5943
NT Start	1449	5102	2493	11883				4714
Contig	GR00689	GR00690	GR00691	GR00720				GR00287 4714
Identification Code Contig	RXA02375	F RXA02378	F RXA02382	RXA02499	RXS02157	RXS02262	RXS02970	F RXA01009
Amino Acid SEQ ID NO	290	294 294	296	298	300	302	304	306
Nucleic Acid	289	293	295	297	299	301	303	305

Enzymes Nucleic Acid SEQ ID NO 309 311 313	of proline Amino Acid SEQ ID NO 308 310 312 314	Enzymes of proline degradation: Nucleic Acid SEQ ID NO 307 Amino Acid 308 Identification Code (Contig.) Contig. 307 308 RXN00023 VV012 310 F RXA00023 GR000 311 312 F RXA02284 GR006 313 314 RXC02498	Contig. VV0127 GR00003 GR00660	NT Start 68158 2 3028	Table 1 (continued) 1 NT Stop Function 64703 CARBOX 454 PROLINE CARBOX 5 CARBOX PROLINE 5 CARBOX PROLINE	Function PROLINE DEHYDROGENASE (EC 1.5.99.8) / DELTA-1- PYRROLINE-5-CARBOXYLATE DEHYDROGENASE (EC 1.5.1.12) PROLINE DEHYDROGENASE (EC 1.5.99.8) / DELTA-1- PYRROLINE-5-CARBOXYLATE DEHYDROGENASE (EC 1.5.12) PROLINE DEHYDROGENASE (EC 1.5.12) PROLINE DEHYDROGENASE (EC 1.5.99.8) / DELTA-1- PYRROLINE-5-CARBOXYLATE DEHYDROGENASE (EC 1.5.1.2) PROTEIN INVOLVED IN PROLINE METABOLISM
			سدو			

Synthesis of 3-Hyo	of 3-Hydoxy-proline:				
	Amino Acid Identification Code Contig	Contig	NT Start	NT Start NT Stop Function	Function
315 316	RXA01491	GR00423 5337		4687	DNA FOR L-PROLINE 3-HYDROXYLASE, COMPLETE CDS
Enzymes of ornith	ornithine, arginine and spermidine metabolism:	nd spei	midine n	netaboli	sm:

Function	GLUTAMATE N.ACETYLTRANSFERASE (EC 2.3.1.35) / AMINO-ACID ACETYLTRANSFERASE (EC 2.3.1.1)	ACETYLGLUTAMATE KINASE (EC 2,7.2.8)	N-ACETYL-GAMMA-GLUTAMYL-PHOSPHATE REDUCTASE (EC 1.2.1.38)	N-ACETYLGLUTAMATE-5-SEMIALDEHYDE DEHYDROGENASE	N-ACETYLGLUTAMATE-5-SEMIALDEHYDE DEHYDROGENASE	ACETYLORNITHINE AMINOTRANSFERASE (EC 2.6.1.11)	ACETYLORNITHINE AMINOTRANSFERASE (EC 2.6.1.11)	ACETYLORNITHINE AMINOTRANSFERASE (EC 2.6.1.11)	ORNITHINE CARBAMOYLTRANSFERASE (EC 2.1.3.3)	ARGININOSUCCINATE SYNTHASE (EC 6.3.4.5)	ARGININOSUCCINATE LYASE (EC 4.3.2.1)	ARGININOSUCCINATE LYASE (EC 4.3.2.1)	ARGININOSUCCINATE LYASE (EC 4.3.2.1)	ORNITHINE CYCLODEAMINASE (EC 4.3.1.12)	SPERMIDINE SYNTHASE (EC 2.5.1.16)	SPERMIDINE SYNTHASE (EC 2.5.1.16)	PUTRESCINE OXIDASE (EC 1.4.3.10)	ARGININE HYDROXIMATE RESISTANCE PROTEIN	N-ACETYL-GAMMA-GLUTAMYL-PHOSPHATE REDUCTASE (EC 1.2.1.38)	CARBAMOYL-PHOSPHATE SYNTHASE SMALL CHAIN (EC 6.3.5.5)	N-ACYL-L-AMINO ACID AMIDOHYDROLASE (EC 3.5 1.14)	N-ACYL-L-AMINO ACID AMIDOHYDROLASE (EC 3.5.1.14)
NT Stop	3076	4075	13327	1536	1826	5251		5943	6224	8116	5253	8962	9611	33436	20230	14190	2142	6743	13037			
NT Start	1913	3125	14106	757	1536	4079		4714	5268	6914	6683	8180	8949	32291	19289	12652	2942	6231	13327			
Contig	GR00640	GR00640	W0122	GR00640	GR00640	GR00640		GR00287	GR00640	GR00640	VV0122	GR00640	GR00640	GR00654	GR00032	GR00424	_	GR00640	VV0122			
Identification Code	RXA02155	RXA02156	RXN02153	F RXA02153	RXA02154	RXA02157	RXS02970	F RXA01009	RXA02158	RXA02160	RXN02162	F RXA02161	F RXA02162	RXA02262	RXA00219	RXA01508	RXA01757	RXA02159	RXN02154	RXS00147	RXS00905	RXS00906
Amino Acid	318	320	322	324	326	328	330	332	334	336	338	340	342	344	346	348	350	352	354	356	358	360
Nucleic Acid	317	319	321	323	325	327	329	331	333	335	337	339	341	343	345	347	349	351	353	355	357	359

ntinued)	Function		N-ACYL-L-AMINO ACID AMIDOHYDROLASE (EC 3.5.1.14)	N-ACYL-L-AMINO ACID AMIDOHYDROLASE (EC 3.5.1.14)	N-ACYL-L-AMINO ACID AMIDOHYDROLASE (EC 3.5.1.14)	CARBAMOYL-PHOSPHATE SYNTHASE LARGE CHAIN (EC 6.3.5.5)	CARBAMOYL-PHOSPHATE SYNTHASE LARGE CHAIN (EC 6.3.5.5)	N-ACYL-L-AMINO ACID AMIDOHYDROLASE (EC 3.5.1.14)	N-ACYL-L-AMINO ACID AMIDOHYDROLASE (EC 3.5.1.14)	
Table 1 (continued)	NT Start NT Stop Function						3198			
⊥a	NT Start						-			
	Contig						GR00654			
	Identification Code Contig		RXS00907	RXS02001	RXS02101	RXS02234	F RXA02234	RXS02565	RXS02937	
	Amino Acid	SEQ ID NO	362	364	366	368	370	372	374	
	_							371		

Histidine metabolism

Function	ATP PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.17)	PHOSPHORIBOSYL-ATP PYROPHOSPHOHYDROLASE (EC 3.6.1.31)	PHOSPHORIBOSYL-AMP CYCLOHYDROLASE (EC 3.5.4.19)	PHOSPHORIBOSYLFORMIMINO-5-AMINOIMIDAZOLE CARBOXAMIDE	RIBOTIDE ISOMERASE (EC 5.3.1.16)	AMIDOTRANSFERASE HISH (EC 2.4.2)	AMIDOTRANSFERASE HISH (EC 2.4.2)	AMIDOTRANSFERASE HISH (EC 2.4.2)	HISF PROTEIN	IMIDAZOLEGLYCEROL-PHOSPHATE DEHYDRATASE (EC 4.2.1.19)	IMIDAZOLEGLYCEROL-PHOSPHATE DEHYDRATASE (EC 4.2.1.19) /	HISTIDINOL-PHOSPHATASE (EC 3.1.3.15)	HISTIDINOL-PHOSPHATE AMINOTRANSFERASE (EC 2.6.1.9)	HISTIDINOL-PHOSPHATE AMINOTRANSFERASE (EC 2.6.1.9)	HISTIDINOL-PHOSPHATE AMINOTRANSFERASE (EC 2.6.1.9)	HISTIDINOL DEHYDROGENASE (EC 1.1.1.23)	PROTEIN INVOLVED IN HISTIDINE METABOLISM	PROTEIN INVOLVED IN HISTIDINE METABOLISM	PROTEIN INVOLVED IN HISTIDINE METABOLISM	MEMBRANE SPANNING PROTEIN INVOLVED IN HISTIDINE METABOLISM
NT Stop	2055	2917	4373	6335		7094	39351	2944	4726	6432	10322		23318	525	10947	12053				
NT Start	2897	3186	4726	7072		7726	39950	2444	5499	7037	10927		24181	4	12044	13378				
Contig	GR00645	GR00645	GR00306	GR00306		GR00306	VV0010	GR00460	GR00306	VV0059	GR00306		VV0112	GR00108	GR00306	GR00306				
Identification Code	RXA02194	RXA02195	RXA01097	RXA01100		RXA01101	RXN01657	F RXA01657	RXAC1098	RXN01104	F RXA01104		RXN00446	F RXA00446	RXAC1105	RXA01106	RXC00930	RXC01096	RXC01656	RXC01158
Amino Acid	376	378	380	382		384	386	388	390	392	394		396	398	400	402	404	406	408	410
Nucleic Acid	375	377	379	381		383	385	387	389	391	393		395	397	399	401	403	405	407	409

Metabolism of aromatic amino acids

Function	3-PHOSPHOSHIKIMATE 1-CARBOXYVINYLTRANSFERASE (EC 2.5.1.19)	4-AMINO-4-DEOXYCHORISMATE LYASE (EC 4)	ANTHRANILATE PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.18)	ANTHRANILATE PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.18)	ANTHRANILATE SYNTHASE COMPONENT I (EC 4.1.3.27)	ANTHRANILATE SYNTHASE COMPONENT I (EC 4.1.3.27)
NT Stop	4345	6948	2577	280	2764	1130
NT Start	3056	5806	3197	က	1211	ဗ
Contig	GR00712	GR00777	VV0247	GR00263	VV0208	GR00264
Identification Code	RXA02458	RXA02790	RXN00954	F RXA00954	RXN00957	F RXA00957
Amino Acid	SEQ ID NO	414	416	418	420	422
Nucleic Acid	SEQ ID NO	413	415	417	419	421

tinued)	Function	CHORISMATE MUTASE (EC 5.4.99.5) / PREPHENATE DEHYDRATASE (EC 4.2.1.51)	CHORISMATE SYNTHASE (EC 4.6.1.4)	CHORISMATE SYNTHASE (EC 4.6.1.4)	INDOLE-3-GLYCEROL PHOSPHATE SYNTHASE (EC 4.1.1.48) INDOLE-3-GLYCEROL PHOSPHATE SYNTHASE (EC 4.1.1.48) / N-(5-PHOSPHO-	RIBOSYL)ANTHRANILATE ISOMERASE (EC 5.3.1.24)	ISOCHORISMATE MUTASE	SHIKIMATE 5-DEHYDROGENASE (EC 1.1.1.25)	OHIKIMATE S-DEHYDROGENASE (EC.1.1.1.25) SHIKIMATE S-DEHYDROGENASE (EC.1.1.1.25)	SHIKIMATE KINASE (EC 2.7.1.71)	TRYPTOPHAN SYNTHASE ALPHA CHAIN (EC 4.2.1.20)	TRYPTOPHAN SYNTHASE BETA CHAIN (EC 4.2.1.20)	TRYPTOPHAN SYNTHASE BETA CHAIN (EC 4.2.1.20)	TYROSINE AMINOTRANSFERASE (EC 2.6.1.5)	PREPHENALE DEHYDROGENAGE (EC 1.3.1.12)	PREPRENATE DEHYDROGENAVE (FC 1.3.1.12)	FREFILENATE DEFITIONOGENAGE (EC. 1.3.1.12) PHOSPHO-2-DEHYDRO-3-DEOXYHEDTONATE ALDOLASE (EC. 4.1.2.15)	PARA-AMINOBENZOATE SYNTHASE COMPONENT I (EC 4.1.3.)	PARA-AMINOBENZOATE SYNTHASE GLUTAMINE AMIDOTRANSFERASE	COMPONENT II (EC 4.1.3) / ANTHRANILATE SYNTHASE COMPONENT II (EC 4.1.3.27)	ANTHRANILATE SYNTHASE COMPONENT II (EC 4.1.3.27)	TRYPTOPHAN SYNTHASE BETA CHAIN (EC 4.2.1.20)	3-OXOADIPATE COA-TRANSFERASE SUBUNIT B (EC 2.8.3.6)	3-OXOADIPATE ENOL-LACTONE HYDROLASE (EC 3.1.1.24) / 4- CARBOXYMUCONOLACTONE	ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1)	ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1)	O-SUCCINYLBENZOIC ACIDCOA LIGASE (EC 6 2.1.26)	I 4-UNITUROAT-2-NAPHTHOATE OCTAFRENTLINANSFERASE (EC. 2.3.7.) 1.4.DIHYDROXY-2-NAPHTHOATE OCTABRENYLTRANSFERASE (EC. 2.3.7.)	HISTIDINOL-PHOSPHATE AMINOTRANSFERASE (EC 2.6.1.9)	HISTIDINOL-PHOSPHATE AMINOTRANSFERASE (EC 2.6.1.9)	ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1)	ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1)	AVPAKTATE AMINOTRANVFERAVE (EC. 2.9.1.1) HISTIDINOT PHOSPHATE AMINOTRANSFERASE (FC. 2.6.1.9)	2-SUCCINYL-6-HYDROXY-2,4-CYCLOHEXADIENE-1-CARBOXYLATE	SYNTHASE / 2-OXOGLUTARATE DECARBOXYLASE (EC 4.1.1.71)	ASPARIATE AMINOTRANSFERASE (EC. 2.0.1.1) NAPHTHOATE SYNTHASE (EC. 4.1.3.36)	O-SUCCINYLBENZOIC ACIDCOA LIGASE (EC 6.2.1.26)	ASPARTATE AMINOTRANSFERASE (EC 2.6.1.1) 3.DEHYDRODI:INATE DEHYDRATASE (EC 4.2.1.10)	טיטרון סהטגטוים ב מרוומיטים אינייישן אינייישן אינייישן אינייישן אינייישן אינייישן אינייישן אינייישן אינייישן א
Table 1 (continued)	NT Stop	12250	12736	991	2821 2007		128	936	1324/ 7795	1553	936	4	3157	3776	32940	668	10260	4087	1753		3778	25887	9889	11099		4		4911	-	525	•	746	38						
<u></u>	NT Start	11306	11507	2	3603 586	8	598	1715	12444 6968	984	26	1140	2027	2499	33858	2	11384	5946	1130		3410	25447	7497	10347		510		4030	2	4		213	408						
-	Contig	GR00754	W0134	GR00477	GR00306 GR00263		GR00795	GR00033	GR00529	GR00477	GR00262	VV0247	GR00263	GR00010	211000	6,400,109	GR00156	GR00156	GR00264		W0208	00000	VV0182	W0182		GR00018		GROOORE		GR00108		GR00163	GK0018						
	Identification Code	RXA02687	RXN01698	F RXA01698	RXA01095 RXA00955		RXA02814	RXA00229	KXA02093 RXA02791	RXA01699	RXA00952	RXN00956	F RXA00956	RXA00064	KXN00448	F KXA00448	F KXA00432	RXA00579	RXA00958		RXN03007	RXN02918	RXN01116	RXN01115	RXS00116	F RXA00116	RXS00391	F RX ADD 393	RXS00446	F RXA00446	RXS00618	F RXA00618	F KXA0062/ RXS01105	RXS02315		RX S02319	RXS02908	RXS03003	270000
	Amino Acid	424	426	428	4 30	į	434	4 36	438	142	. 444	446	448	450	452	454 466	4.78 8.78	460	462		464	466	468	470	472	474	476	4 4 0 4 0 4	482	484	486	488	064 064	494	904	4.98 4.98	200	502 504	ř
	Nucleic Acid	423	425	427	429 4 31	· •	433	435	43/ 439	441	443	445	447	449	451	453 466	457	459	461		463	465	467	469	471	473	475	479	481	483	485	487	489 491	493	90	495 497	499	501 503	3

Table 1 (continued)	NT Start NT Stop Function	S-ADENOSYLMETHIONINE 2-DEMETHYLMENAQUINONE	METHYLTRANSFERASE (EC 2.1)	MEMBRANE SPANNING PROTEIN INVOLVED IN METABOLISM OF AROMATIC	AMINO ACIDS AND RIBOFLAVIN	MEMBRANE SPANNING PROTEIN INVOLVED IN METABOLISM OF AROMATIC	AMINO ACIDS	CYTOSOLIC PROTEIN INVOLVED IN METABOLISM OF AROMATIC AMINO	ACIDS	MEMBRANE SPANNING PROTEIN INVOLVED IN METABOLISM OF AROMATIC	AMINO ACIDS
	Identification Code Contig.	RXS03074		RXC01434		RXC02080		RXC02789		RXC02295	
	Amino Acid SEO ID NO	206		508		510		512		514	
	Nucleic Acid SEQ ID NO	505		207		609		511		513	

Aminobutyrate metabolism

Function	4-aminobutyrate aminotransferase (EC 2.6.1.19)	ACETYLORNITHINE AMINOTRANSFERASE (EC 2.6.1.11)	ACETYLORNITHINE AMINOTRANSFERASE (EC 2.6.1.11)
NT Stop	1697	6081	5943
NT Start	999	4/14	4714
Contig	VV0035	VV0021	GR00287
Identification Code	RXN03063	KXNC2970	F RXA01009
Amino Acid SEQ ID NO	516	518	520
Nucleic Acid SEQ ID NO			

Vitamins, vitamin-like substances (cofactors), nutraceuticals

Thiamine metabolism

Function	THIAMIN BIOSYNTHESIS PROTEIN THIC	THIAMIN-MONOPHOSPHATE KINASE (EC 2.7.4.16)	THIAMIN-PHOSPHATE PYROPHOSPHORYLASE (EC 2.5.1.3)	THIF PROTEIN	THIG PROTEIN	THIG PROTEIN	HYDROXYETHYLTHIAZOLE KINASE (EC 2.7.1.50)	APBA PROTEIN	THIAMIN BIOSYNTHESIS PROTEIN X	PHOSPHOMETHYL PYRIMIDINE KINASE (EC 2.7.4.7)	PHOSPHOMETHYLPYRIMIDINE KINASE (EC 2 7 4 7)	PYRIDOXINE KINASE (EC 2.7.1.35)	CYTOSOLIC KINASE INVOLVED IN METABOLISM OF SUGARS AND THIAMIN			
NT Stop	4819	995	4	2286	4	378	1032	633	2557	2446	2446	27905	22858	616		
NT Start	2945	9	609	3206	162	983	229	1532	1988	1019	1019	27306	22187	2		
Contig	GR00431	GR00291	GR00393	GR00403	GR00394	GR00394	GR00348	GR00227	GR00699	VV0270	GR00348	00000	050000	GR00451		
Identification Code	RXA01551	RXA01019	RXA01352	RXA01381	RXA01360	RXA01361	RXA01208	RXA00838	RXA02400	RXN01209	F RXA01209	RXN01413	RXN01617	F RXA01617	RXS01807	RXC01021
Amino Acid SEQ ID NO	522	524	526	528	530	532	534	536	538	540	542	544	546	548	550	552
Nucleic Acid				527												

PYRIDOXINE KINASE (EC 2.7.1.35), pyridoxal/pyridoxine/pyridoxamine kinase

Function

NT Stop 7077

NT Start 7868

Identification Code

Contig GR00509

RXA01807

Amino Acid SEQ ID NO 596

Nucleic Acid SEQ ID NO 595

	NT Start NT Ston Function
vin metabolism	
vin metabo	Nucleic Acid Amino Acid Identification Code Contin
. <u>></u>	Amin Aria
Ribofla	Nicolar Acid

art NT Stop Function	5371 diaminohydroxyphosphoribosylaminopyrimidine deaminase (EC 3.5.4.26) / 5-amino-	15282	7286	17197	7777	17688		18356	2388 RIBOFLAVIN KINASE (EC 2.7.1.26) / FMN ADENYLYLTRANSFERASE (EC	2.7.7.2)	1736 NICOTINATE-NUCLEOTIDEDIMETHYLBENZIMIDAZOLE	2388 RIBOFLAVIN KINASE (EC 2.7.1.26) / FMN ADENYLYLTRANSFERASE (EC	2.7.7.2)			629			MEMBRANE SPANNING PROTEIN INVOLVED IN RIBOFLAVIN METABOLISM	PROTEIN INVOLVED IN RIBOFLAVIN METABOLISM	56 Predicted nucleotidyltransferases	CYTOSOLIC PROTEIN INVOLVED IN METABOLISM OF RIBOFLAVIN AND	CIPIOS	MEMBRANE SPANNING PROTEIN INVOLVED IN METABOLISM OF AROMATIC AMINO ACIDS AND RIBOFLAVIN		
NT Start	4388	14299	_	 		17212	_	17778	3410		2809	3410		8993		_	767	1363			602					
Contig	VV0130	GR00654	W0130	GR00654	W0130	GR00654		GR00654	GR00423		GR00639	GR00423		W0191	GR00484	W0213	VV0319	VV0109			GR00691					
Identification Code	RXN02246	F RXA02246 RXA02247	RXN02248	F RXA02248	RXN02249	F RXA02249		RXA02250	RXA01489		RXA02135	RXA01489		RXN01712	F RXA01712	RXN02384	RXN01560	RXN00667	RXC01711	RXC02380	F RXA02380	RXC02921		RXC01434	lism	
Amino Acid	554 554	556 558	560	562	564	999		568	570		572	574		576	578	580	582	584	586	588	290	592		594	Vitamin B6 metabolism	
Nucleic Acid	553	555 557	559	561	563	565		267	999		571	573		575	577	579	581	583	585	587	589	591		593	Vitamin E	

Table 1 (continued)	d), nicotinamide, NAD and NADP
	Nicotinate (nicotinic acid),

Function	NICOTINATE PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.11)	NICOTINATE PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.11)	NICOTINATE PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.11)	NICOTINATE-NUCLEOTIDE PYROPHOSPHORYLASE (CARBOXYLATING) (EC	2.4.2.19) QUINOLINATE SYNTHETASE A
NT Start NT Stop	23901	4	488	6436	5593
NT Start	22564	774	3	2600	4310
Contig	VV0084	GR00701	GR00766	GR00632	GR00632
Identification Code	RXN02754	F RXA02405	F RXA02754	RXA02112	RXA02111
Amino Acid	598	009	602	604	909
Nucleic Acid	597	599	601	603	605

NAD Biosynthesis

Function	NH(3)-DEPENDENT NAD(+) SYNTHETASE (EC 6.3.5.1)	NICOTINATE PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.11)
NT Start NT Stop	2104	23901
NT Start	0 1274	22564
Contig	GR00300	VV0084 2
Identification Code Contig.	RXA01073	RXN02754
Amino Acid SEQ ID NO	809	610
Nucleic Acid SEQ ID NO	209	609

Pantothenate and Coenzyme A (CoA) biosynthesis

Function	ASPARTATE 1-DECARBOXYLASE PRECURSOR (EC 4.1.1.11)	PANTOATEBETA-ALANINE LIGASE (EC 6.3.2.1)	3-METHYL-2-OXOBUTANOATE HYDROXYMETHYLTRANSFERASE (EC 2.1.2.11)	/ DECARBOXYLASE (EC 4.1.1.44)	3-METHYL-2-OXOBUTANOATE HYDROXYMETHYLTRANSFERASE (EC 2.1.2.11)	PANTOATEBETA-ALANINE LIGASE (EC 6.3.2.1)	KETOL-ACID REDUCTOISOMERASE (EC 1.1.1.86)	KETOL-ACID REDUCTOISOMERASE (EC 1.1.1.86)	DNA/PANTOTHENATE METABOLISM FLAVOPROTEIN	PANTOTHENATE KINASE (EC 2.7.1.33)	2-DEHYDROPANTOATE 2-REDUCTASE (EC 1.1.1.169)	PROTEIN INVOLVED IN METABOLISM OF S-ADENOSYLMETHIONINE, PURINES	AND PANTOTHENATE
NT Stop	10859	1121	48402		1960	25964		1530	7049	8540			
NT Start	10452	1957	47590		2766	25167		1075	5784	7572			
Contig.	GR00662	GR00555	VV0127		GR00555	GR00424		GR00321	GR00654	GR00156			
Identification Code	RXA02299	RXA01928	RXN01929		F RXA01929	RXA01521	RXS01145	F RXA01145	RXA02239	RXA00581	RXS00838	RXC02238	
Amino Acid SEQ ID NO	612	614	616		618	620	622	624	979	628	630	632	
Nucleic Acid	611	613	615		617	619	621	623	625	627	629	631	

Biotin metabolism

BIOTIN SYNTHESIS PROTEIN BIOC	8754	8272	VV0028	RXN03058	634	633	
					SEO ID NO	SEQ ID NO	
Function	NT Stop	NT Start	Contig.	Identification Code	Amino Acid	Nucleic Acid	

tinued) Function	BIOTIN SYNTHESIS PROTEIN BIOC BIOTIN SYNTHESIS PROTEIN BIOC ADENOSYLMETHIONINE-8-AMINO-7-OXONONANOATE AMINOTRANSFERASE	DETHIOBILIN SYNTHETASE (EC 6.3.3.3) BIOTIN SYNTHASE (EC 2.8.1.6) NIFS PROTEIN NIFS PROTEIN NIFS PROTEIN NIFS PROTEIN NIFS PROTEIN	NIFS PROTEIN NIFS PROTEIN NIFU PROTEIN		Function	LIPOIC ACID SYNTHETASE LIPOATE-PROTEIN LIGASE B (EC 6) LIPOATE-PROTEIN LIGASE A (EC 6) DIMYDROLIPOAMIDE SUCCINYLTRANSFERASE COMPONENT (E2) OF 2- COCKI ITARATE DEHYDROGENASE COMPILES 3 1 61)	LIPOAMIDE DEHYDROGENASE COMPONENT (E3) OF BRANCHED-CHAIN AI PHA-KETO ACID DEHYDROGENASE COMPI FX (FC 1 8 1 4)	LIPOAMIDE DEHYDROGENASE COMPONENT (E3) OF BRANCHED-CHAIN ALPHA-KETO ACID DEHYDROGENASE COMPLEX (EC 1.8.1.4)		Function	5.10-METHYLENETETRAHYDROFOLATE REDUCTASE (EC 17.99 5) 5-FORMYLTETRAHYDROFOLATE CYCLO-LIGASE (EC 6.3.3.2) 5-FORMYLTETRAHYDROFOLATE CYCLO-LIGASE (EC 6.3.3.2)	DIHYDROFOLATE REDUCTASE (EC 1.5.1.3) FORMYLTETRAHYDROFOLATE DEFORMYLASE (EC 3.5.1.10) FORMYLTETRAHYDROFOLATE DEFORMYLASE (EC 3.5.1.10)	METHYLENETETRAHYDROFOLATE DEHYDROGENASE (EC 1.5.1.5) / METHENYLTETRAHYDROFOLATE CYCLOHYDROLASE (EC 3.5.4.9)	GTP CYCLOHYDROLASE I (EC 3.5.4.16) DIHYDRONEOPTERIN ALDOLASE (EC 4.1.2.25)
Table 1 (continued)	12014 4309 2288	1610 4408 22879 15608 897 11209	2949 4 2986 3435		NT Stop	3549 2366 1527				NT Stop	17400 1003 6	1792 4 9788 559	1279	21509 22749
Ta	11532 3650 3556	2281 3407 23967 16681 79 10037	3563 438 1724 2989		NT Start	2506 1614 472				NT Start	18281 503 500	17469 8868 23	428	20922 22360
Contig	GR10040 GR00025 GR00166	GR00166 GR00047 GR00032 VV0123 GR00040	GR00100 GR00782 GR00723 GR00723		Contig	GR00495 GR00495 GR00632			*	Contig	GR00758 VV0296 GR00616	GR00014 VV0082 GR00384	GR00116	GR00424 GR00424
Identification Code	F RXA02903 RXA00166 RXA00633	RXA00632 RXA00295 RXA00223 RXN00262 F RXA00262 RXN00435	F RXA00435 F RXA02801 RXA02516 RXA02517		Identification Code	RXA01747 RXA01746 RXA02106 RXS01183	RXS01260	RXS01261	<u>.s</u>	Identification Code	RXA02717 RXN02027 F RXA02027	RXA00106 RXN01321 F RXA01321	RXA00461	RXA01514 RXA01516
Amino Acid SEQ ID NO	636 638 640	642 644 646 648 650	654 656 658 660	pic	Amino Acid	664 664 666 668	670	672	Folate biosynthesis	Amino Acid	674 676 678	680 682 684	686	688 690
Nucleic Acid SEQ ID NO	635 63 <i>7</i> 639	641 643 645 647 649	653 655 657 659	Lipoic Acid	Nucleic Acid	661 663 665 667	699	671	Folate bi	Nucleic Acid	673 675 677	679 681 683	685	687 689

ntinued)	Function	DIHYDROPTEROATE SYNTHASE (EC 2.5.1.15)	DIHYDROPTEROATE SYNTHASE (EC 2.5.1.15)	DIHYDROFOLATE REDUCTASE (EC 1.5.1.3)	FOLYLPOLYGLUTAMATE SYNTHASE (EC 6.3.2.17)	2-AMINO-4-HYDROXY-6-HYDROXYMETHYLDIHYDROPTERIDINE	PYROPHOSPHOKINASE (EC 2.7.6.3)	PARA-AMINOBENZOATE SYNTHASE COMPONENT I (EC 4.1.3)	PARA-AMINOBENZOATE SYNTHASE GLUTAMINE AMIDOTRANSFERASE	COMPONENT II (EC 4.1.3) / ANTHRANILATE SYNTHASE COMPONENT II (EC	4.1.3.27) 4.4MINO.4.DEOXYCHORISMATETVASE (EC.4)	DIHYDROFOLATE REDUCTASE (FC 1.5.1.3)	5-METHYLTETRAHYDROFOLATEHOMOCYSTEINE METHYLTRANSFERASE	(EC 2.1.1.13)	5-METHYLTETRAHYDROFOLATEHOMOCYSTEINE METHYLTRANSFERASE	(EC 2.1.1.13)	5-METHYLTETRAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTEINE	METHYLTRANSFERASE	5-METHYLTETRAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTEINE	METHYLTRANSFERASE (EC 2.1.1.14)	5-METHYLTETRAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTEINE	METHYLIKANSFEKASE (EC. 2.1.1.14)	5-WETHYLTETRAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTEINE METHYLTBANSFEBASE /FC 2 4 1 1 1/1	S-METHYLTETRAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTFINE	METHYLTRANSFERASE (EC 2.1.1.14)	5-METHYLTETRAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTEINE	METHYLTRANSFERASE (EC 2.1.1.14)	5-METHYLTETRAHYDROFOLATEHOMOCYSTEINE METHYLTRANSFERASE	(EC 2.1.1.13)	PROTEIN INVOLVED IN FOLATE METABOLISM MEMBRANE SPANNING PROTEIN INVOLVED IN FOLATE METABOLISM	ATP-BINDING PROTEIN INVOLVED IN FOLATE METABOLISM
Table 1 (continued)	NT Stop	22364	4784	17924	1371	23228		4087	1753		8048	17924	11726		9		10717		5295		5731			4730		15447					
-	NT Start	21513	4026	17469	2903	22752		5946	1130		5806	17469	9228		2483		8483		3496		5252			5254		14764					
	Contig	GR00424	GR00613	GR00014	GR00280	GR00424		GR00156	GR00264		GB00777	GR00014	VV0302		GR00646		VV0126		GR00629		GR00629			GR00751		GR00752					
	Identification Code	RXA01515	RXA02024	RXA00106	RXA00989	RXA01517		RXA00579	RXA00958		RXA02790	RXA00106	RXN02198		F RXA02198		RXN02085		F RXA02085		F RXA02086		KXN02648	F RXA02648		F RXA02658		RXS02197		RXC00988	RXC01942
	Amino Acid SEQ ID NO	692	694	969	869	200		702	704		706	208	710		712		714		716		718	ć	07)	722		724		726		728	732
	Nucleic Acid	691	693	695	269	669		701	703		705	202	402		711		713		715		717		817	721		723		725		727	731

Molybdopterin Metabolism

Function	MOLYBDOPTERIN BIOSYNTHESIS MOEB PROTEIN	MOLYBDOPTERIN BIOSYNTHESIS MOEB PROTEIN	MOLYBDOPTERIN BIOSYNTHESIS MOEB PROTEIN	MOLYBDOPTERIN (MPT) CONVERTING FACTOR, SUBUNIT 2	MOLYBDOPTERIN (MPT) CONVERTING FACTOR, SUBUNIT 2	MOLYBDOPTERIN CO-FACTOR SYNTHESIS PROTEIN	MOLYBDOPTERIN CO-FACTOR SYNTHESIS PROTEIN	MOLYBDOPTERIN CO-FACTOR SYNTHESIS PROTEIN
NT Start NT Stop	16299	474	796	17369	362	18275	196	1087
NT Start	17369	7	362	17824	6	18742	2	830
Contig	VV0112	GR00783	GR00103	VV0112	GR00103	W0112	GR00104	GR00105
Identification Code	RXN02802	F RXA02802	F RXA00438	RXN00437	F RXA00437	RXN00439	F RXA00439	F RXA00442
Amino Acid SEQ ID NO	734	736	738	740	742	744	746	748
Nucleic Acid SEQ ID NO								

Table 1 (continued)	Function	MOLYBDENUM COFACTOR BIOSYNTHESIS PROTEIN CB	MOLYBDOPTERIN CO-FACTOR SYNTHESIS PROTEIN	MOLYBDOPTERIN CO-FACTOR SYNTHESIS PROTEIN	5-METHYLTETRAHYDROPTEROYLTRIGLUTAMATEHOMOCYSTEINE	METHYLTRANSFERASE (EC 2.1.1.14)	DIHYDRONEOPTERIN ALDOLASE (EC 4.1.2.25)	DIHYDROPTEROATE SYNTHASE (EC 2.5.1.15)	DIHYDROPTEROATE SYNTHASE (EC 2.5.1.15)	MOLYBDOPTERIN-GUANINE DINUCLEOTIDE BIOSYNTHESIS PROTEIN A	MOLYBDOPTERIN BIOSYNTHESIS MOEA PROTEIN	MOLYBDOPTERIN BIOSYNTHESIS MOEA PROTEIN	MOLYBDOPTERIN BIOSYNTHESIS MOEA PROTEIN	MOLYBDOPTERIN BIOSYNTHESIS CNX1 PROTEIN	(D90909) pterin-4a-carbinolamine dehydratase [Synechocystis sp.]	2-AMINO-4-HYDROXY-6-HYDROXYMETHYLDIHYDROPTERIDINE DYDDDHOSDHOKINASE (FC 2.7.6.3)	MINIOR TOOL TOOL TOOL (FO 2.1.0.3)	FLAVOHEMOPROTEIN / DIHYDROPTERIDINE REDUCTASE (FC 16 99 7)	OXYGEN-INSENSITIVE NAD(P)H NITROREDUCTASE (EC. 1) /	DIHYDROPTERIDINE REDUCTASE (EC1.6.99.7)										
ble 1 (co	NT Stop	654	18779	793			5295		5731				4730		15447		22749	22364	4784	704	1268		1207	069	3965	23228	7077	5		
<u>'</u>	NT Start	196	19942	2	_		3496		5252				5254		14764		22360	21513	4026	1264	2476		2	1274	9684	22752	7440) †		
	Contig	GR00104	W0112	GR00105			GR00629		GR00629				GR00751		GR00752		GR00424	GR00424	GR00613	GR00488	GR00488		GR00568	GR00748	GR00665	GR00424	87.0707	2		
	Identification Code	RXA00440	RXN00441	F RXA00441	RXN02085		F RXA02085		F RXA02086		RXN02648		F RXA02648		F RXA02658		RXA01516	RXA01515	RXA02024	RXA01719	RXA01720	RXS03223	F RXA01970	RXA02629	RXA02318	RXA01517	DVN01304	RXS02556	RXS02560	
	Amino Acid	750	752	754	756		758		760		762		764		766		768	770	772	774	776	778	780	782	784	786	788	790	792	!
	Nucleic Acid	749	751	753	755		757		759		761		763		765		191	692	771	773	775	777	779	781	783	785	787	780	791	

Vitamin B₁₂, porphyrins and heme metabolism

Function	GLUTAMATE-1-SEMIALDEHYDE 2,1-AMINOMUTASE (EC 5.4.3.8)	FERROCHELATASE (EC 4.99.1.1)	FERROCHELATASE (EC 4.99.1.1)	HEMK PROTEIN	OXYGEN-INDEPENDENT COPROPORPHYRINOGEN III OXIDASE (EC 1)	PORPHOBILINOGEN DEAMINASE (EC 4.3.1.8)	PORPHOBILINOGEN DEAMINASE (EC 4.3.1.8)	UROPORPHYRINOGEN DECARBOXYLASE (EC 4.1.1.37)	PORPHOBILINOGEN DEAMINASE (EC 4.3.1.8)	PORPHOBILINOGEN DEAMINASE (EC 4.3.1.8)
NT Stop	1451	9400	8596	1274	11276	22854	17340	306	23362	17816
NT Start	2752	10509	7910	2206	10137	22456	16906	1427	22805	17379
Contig	GR00082	GR00023	GR00163	GR00051	GR00242	VV0007	GR00720	GR00081	VV0007	GR00720
Identification Code	RXA00382	RXA00156	RXA00624	RXA00306	RXA00884	RXN02503	F RXA02503	RXA00377	RXN02504	F RXA02504
Amino Acid SEQ ID NO	794	962	798	800	802	804	806	808	810	812
Nucleic Acid SEQ ID NO	793	795	797	799	801	803	805	807	809	811

ntinued)	Function	PRECORRIN-6Y METHYLASE (EC 2.1.1)	PRECORRIN-6Y METHYLASE (EC 2.1.1.)	UROPORPHYRIN-III C-METHYLTRANSFERASE (EC 2.1.1.107)	UROPORPHYRIN-III C-METHYLTRANSFERASE (EC 2.1.1.107) / UROPORPHYRINOGEN-III SYNTHASE (EC 4.2.1.75)	UROPORPHYRIN-III C-METHYLTRANSFERASE (EC 2.1.1.107) /	ONCOUNTY IN INCOME THE THE THE THE THE THE THE THE THE TH	UROPORPHYRINOGEN-III SYNTHASE (EC 4.2.1.75)	PROTOPORPHYRINOGEN OXIDASE (EC 1.3.3.4)	PROTOPORPHYRINOGEN OXIDASE (EC 1.3.3.4)	PROTOPORPHYRINOGEN OXIDASE (EC 1.3.3.4)	COBYRIC ACID SYNTHASE	COBALAMIN (5'-PHOSPHATE) SYNTHASE	NICOTINATE-NUCLEOTIDEDIMETHYLBENZIMIDAZOLE	PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.21)	COBINAMIDE KINASE / COBINAMIDE PHOSPHATE GUANYLYLTRANSFERASE	COBG PROTEIN (EC 1)	HEMIN-BINDING PERIPLASMIC PROTEIN HMUT PRECURSOR	HEMK PROTEIN	HEMK PROTEIN	CYTOSOLIC PROTEIN INVOLVED IN PORPHYRIN METABOLISM		Function		L-GUIDACHONE CATALOGUE (EC. 1-13.8)	L-GUCUNOLACTIONS OXIDASE (EC. 1.13.8)	L-GUILLONGLACTIONS (FC 1.13.8)	2.9-UNETION-CONIC ACID REDUCTABLE (EC. 1.7.1)	ZIGHONDELIOONIC ACID REDUICTARE (EC. 1)	oxodilitarate comialdebude debudronanse (FC 1.7.1)	OXOQUARINE SERINGUING CENTY OXOGUAS (LC 1.2.1.7) ACETOACETYI COA REDITOTASE (EC 1.1.3)	ACCUMENTATION OF THE CONTRACT OF THE CONTRACT OF STREET	OXIDOREDUCTASE INVOLVED IN METABOLISM OF VITAMIN C PRECURSORS		Function	O ADENOAM METHIONINE O DEMETHAL MENADLENONE	METHYLTRANSFERASE (EC 2.1)	
Table 1 (continued)	NT Stop	524	4 1	749	5973	9	274	-	2863	9	2863	1787	801	1736		2841	552	663					NT Stop	970	040	541	2258	38/2	1339 828	0.50					NT Stop			
Ĥ	NT Start	1849	1248	1498	4180	929	1103	701-	4206	287	3876	2536	1721	2809		3362	-	1739					NT Start	7544	1107	7	1/3/	8/04	1540	2					NT Start			
	Contig	00008	GR00330	GR004/4	VV0226	GR00078	020000	6 20010	VV0223	GR00081	GR00082	GR00365	GR00639	GR00639		GR00639	VV0088	VV0082					Contig.		211000	GK00036	CK00097	VV0005	CBL00183	20000					Contig			
	Identification Code	RXN01162	F RXA01162	KXAU1692	KXN003/1	F RXA00371	E BY A 00274	100000	RXN00383	F RXA00376	F RXA00383	RXA01253	RXA02134	RXA02135		RXA02136	RXN03114	RXN01810	RXS03205	F RXA00306	RXC01715	ors	Identification Code	007001450	RAIN00420	F KXA00420	F KXA00426	KANUU/U8	PX 402373	RX S00389	RX S00419	RXC00416	RXC02206		Identification Code	PX S03074	100000	
	Amino Acid SEQ ID NO	814	816	818	028	822	70	†	826	828	830	832	834	836		838	840	842	844	846	848	C precursors	Amino Acid		000	258	854	856	850 850	862	864	866	868	5	Amino Acid	25 LO 140	2	
	Nucleic Acid SEQ ID NO	813	815	817	819	821	673	0.50	825	827	829	831	833	835		837	839	841	843	845	847	Vitamin C	Nucleic Acid		9 4 5	851	853	855	050	861	863	865	867	Vitamin K2	Nucleic Acid	869	5	

Function	S-ADENOSYLMETHIONINE:2-DEMETHYLMENAQUINONE	MEINTLIANDPERADE (EU Z.1.;) 2-SUCCINYL-6-HYDROXY-2-4-CYCLOHEXADIENE-1-CARBOXYLATE 2-SUCCINYL-6-HYDROXY-2-4-7-7-1-1-1-1-1-1-1-1-1-1-1-1-1-1-1-1-1	STNINASEZ-CONOGLOTARSTE DECAMBONITASE (EC. 4.1.1.7.7.) 1 APHTHOATE SYNTHASE (EC. 4.1.3.36) 1 APHTHOATE SYNTHASE (EC. 4.1.3.36)	1,4-DIHYDROX1-2-NAPHTHOATE OCTAFRENTLINANSFERASE (EC 2.5) 1,4-DIHYDROXY-2-NAPHTHOATE OCTAFRENYLTRANSFERASE (EC 2.5) 0-SUCCINYLBENZOIC ACIDCOA LIGASE (EC 6.2.1.26) 0-SUCCINYLBENZOIC ACIDCOA LIGASE (EC 6.2.1.26)		Function	3-DEMETHYLUBIQUINONE-9 3-METHYLTRANSFERASE (EC 2.1.1.64) 3-DEMETHYLUBIQUINONE-9 3-METHYLTRANSFERASE (EC 2.1.1.64) 3-DEMETHYLUBIQUINONE-9 3-METHYLTRANSFERASE (EC 2.1.1.64) UBIQUINONE/MENAQUINONE BIOSYNTHESIS METHLYTRANSFERASE UBIE	(EC 2.1.1-) COMA OPERON PROTEIN 2		ays			Function	RIBOSE-PHOSPHATE PYROPHOSPHOKINASE, PRPP synthetase (EC 2.7 6.1) AMIDOPHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.14)	AMILOPHUSPHURIBOSTLINANSFERASE (EC. 2.4.2.1.4) PHOSPHORIBOSYLAMINEGLYCINE LIGASE (EC. 6.3.4.13) PHOSPHORIBOSYLAMINE-GLYCINE LIGASE (EC. 6.3.4.13)	PHOSPHORIBOSYLAMINEGLYCINE LIGASE, GARS (EC 6.3.4.13) PHOSPHORIBOSYLAMINEGLYCINE LIGASE (EC 6.3.4.13) / PHOSPHORIBOSYLFORMYLGLYCINAMIDINE CYCLO-LIGASE (EC 6.3.3.1) /	PHOSPHORIBOSYLGLYCINAMIDE FORMYLTRANSFERASE (EC 2.1.2.2) PHOSPHORIBOSYLGLYCINAMIDE FORMYLTRANSFERASE 2 (EC 2.1.2)	
Table 1 (continued)	645	6383	10933	4911 2750		NT Stop	1808 249 2384 12547		Nucleotides	dine biosynthesis pathways			NT Stop	213 9581	501 10362 1713	780 4285	9054	
NT Start	1142	8011	2266	4030		NT Start	2389 986 3073 13299		•	synthes			NT Start	1187 8235	61 11624 1450	4875	10277	
Contig.	GR10044	GR00665	GR00665	GR00086 GR00086		Contig	GR00283 GR00642 GR00665 VV0135		nd othe	dine bio			Contig	GR00352 W0103	GR00148 VV0135 GR00165	GR00164 GR00746	GR00418	
Identification Code	F RXA02906	RXA02315	RXA02319	F RXA00393 F RXA00391 RXA00391 RXS02908	nthesis	Identification Code	RXA00997 RXA02189 RXA02311 RXN02912	RXS00998	Pyrimidines an	Regulation of purine and pyrimi	_	10	Identification Code	RXA01215 RXN00558	F KXAUU558 RXN00626 F RXA00629	F RXA00626 RXA02623	RXA01442	
Amino Acid	872	874	876	880 882 884 884	Ubiquinone biosynthesis	Amino Acid	SEQ ID NO 886 888 890 892	894	and	ion of puri	Purine metabolism	Purine Biosynthesis	Amino Acid	896 898 898	900 902 90 4	906 908	910	
Nucleic Acid	871	873	875	879 881 883	Ubiquin	Nucleic Acid	SEQ ID NO 885 887 889 891	893	Purines	Regulat	Purine r	Purine E	Nucleic Acid	895 897 897	899 901 903	905 907	606	

Table 1 (continued)	Function	PHOSPHORIBOSYLFORMYLGLYCINAMIDINE SYNTHASE (EC 6.3.5.3)	PHOSPHORIBOSYLAMIDOIMIDAZOLE-SUCCINOCARBOXAMIDE SYNTHASE	(EC 6.3.2.6)	PHOSPHORIBOST L'TORINT L'OLL CONSTITUTION DE L'ACTORITATION DE L'A	PHOSPHORIBOSYLFORMYLGLYCINAMIDINE CYCLO-LIGASE (EC 6.3.3.1)	PHOSPHORIBOSYLFORMYLGLYCINAMIDINE CYCLO-LIGASE (EC 6.3.3.1)	PHOSPHORIBOSYLAMINOIMIDAZOLE CARBOXYLASE ATPASE SUBUNIT (EC	4.1.21)	PHOSPHORIBOSYLAMINOIMIDAZOLE CARBOXYLASE ATPASE SUBUNIT (EC	4.1.1.21)	PHOSPHORIBOSYLAMINOIMIDAZOLE CARBOXYLASE CATALYTIC SUBUNIT	(EC 4.1.1.21)	PHOSPHORIBOSYLAMINOIMIDAZOLE CARBOXYLASE CATALYTIC SUBUNIT	(EC 4.1.1.21)	PHOSPHORIBOSYLAMINOIMIDAZOLE CARBOXYLASE CATALYTIC SUBUNIT	(EC 4.1.1.21)	PHOSPHORIBOSYLAMINOIMIDAZOLE CARBOXYLASE (EC 4.1.1.21)	ADENYLOSUCCINA IE LYASE (EC 4.3.2.2)	PHOSPHORIBOSYLAMINOIMIDAZOLECARBOXAMIDE FORMITLIRANSFERASE (EC 2.1.2.3) / IMP CYCLOHYDROLASE (EC 3.5.4.10)				
able 1 (cc	NT Stop	5636	638	269	280	2937	3939		10/83	818	7495	5984		725		8863		5		911		1373	2220	2715
F	NT Start	3351	54	23	2	2269	3049		9614	15	7809	4788		1534		8369		127		1120		498	793	4274
	Contig	VV0103	GR00786	GR00138	GR00150	GR00139	GR00163		VV0103	GR00147	GR00204	VV0078		GR00676		VV0078		GR00677		GR00678		GR00304	GR00163	GR00746
	Identification Code	RXN00537	F RXA02805	F RXA00537	F RXA00561	RXA00541	RXA00620		RXN00770	F RXA00557	F RXA00770	RXN02345		F RXA02345		RXN02350		F RXA02346		F RXA02350		RXA01087	RXA00619	RXA02622
	Amino Acid	912	914	916	918	920	922		924	926	928	930		932		934		936		938		940	942	944
	Nucleic Acid	911	913	915	917	919	921		923	925	927	626	2	931		933		935	}	937	}	939	941	943

GMP, GDP, AMP and ADP synthesis, from inosine-5'-monophosphate (IMP)

	INOSINE-5'-MONOPHOSPHATE DEHYDROGENASE (EC 1.1.1.205)	NOSINE-5'-MONOPHOSPHATE DEHYDROGENASE (EC. 1.1.1.205)	INOSINE-5'-MONOPHOSPHATE DEHYDROGENASE (EU T. 1. 203)	INOSINE-5'-MONOPHOSPHATE DEHYDROGENASE (EC 1.1.1.205)	GMP SYNTHASE [GLUTAMINE-HYDROLYZING] (EC 6.3.5.2)	GMP SYNTHASE (EC 6.3.4.1)	SUANYLATE KINASE (EC 2.7.4.8)	ADENYLOSUCCINATE SYNTHETASE (EC 6.3.4.4)	ADENYLOSUCCINATE LYASE (EC 4.3.2.2)	ADENYLATE KINASE (EC 2.7.4.3)	NUCLEOSIDE DIPHOSPHATE KINASE (EC 2.7.4.5)
Function	INOSINE-	INOSINE-	INOSINE-	INOSINE	GMP SYN	GMP SYN	GUANYLA	ADENYLO	ADENYLO	ADENYLA	NUCLEOS
NT Start NT Stop	20583	1644	534	497	25302	2097	5146	16476	2220	10985	3362
NT Start	19066	1171	-	1927	23734	712	4577	17765	793	10443	3769
Contig	00000	GR00122	GR00121	GR00715	00000	GR00120	GR00654	GR00418	GR00163	GR00179	GR00040
Identification Code	RXN00488	F RXA00492	F RXA00488	RXA02469	RXN00487	F RXA00487	RXA02237	RXA01446	RXA00619	RXA00688	RXA00266
Amino Acid	946	948	950	952	954	926	958	096	362	964	996
Nucleic Acid	945	947	949	951	953	955	957	959	961	963	965

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Table 1 (continued)

GMP REDUCTASE (EC 1.6.6.8)	AMP NUCLEOSIDASE (EC 3.2.2.4)	AMP NUCLEOSIDASE (EC 3.2.2.4)
1775	3323	34
_		
GR00121	VV0152	GR00659
RXA00489 GR00121		
	RXN02281	F RXA02281
	654 1775	_

Pyrimidine metabolism

Pyrimidine biosynthesis de novo:

Function	CARBAMOYL-PHOSPHATE SYNTHASE SMALL CHAIN (EC 6.3.5.5)	ASPARTATE CARBAMOYLTRANSFERASE CATALYTIC CHAIN (EC 2.1.3.2)	DIHYDROOROTASE (EC 3.5.2.3)	DIHYDROOROTATE DEHYDROGENASE 'F' 1.3.3.1)	OROTATE PHOSPHORIBOSYLTRANSFEK, SE (EC 2 4.2.10)	OROTIDINE 5: PHOSPHATE DECARBOXYLASE (EC 4.1.1.23)	URIDYLATE KINASE (EC 2.7.4)	URIDYLATE KINASE (EC 2.7.4)	THYMIDYLATE SYNTHASE (EC 2.1.1.45)	THYMIDYLATE KINASE (EC 2.7.4.9)	NUCLEOSIDE DIPHOSPHATE KINASE (EC 2.7.4.6)	CYTIDYLATE KINASE (EC 2.7.4.14)	CTP SYNTHASE (EC 6.3.4.2)	CARBAMOYL-PHOSPHATE SYNTHASE LARGE CHAIN (EC 6.3.5.5)	CARBAMOYL-PHOSPHATE SYNTHASE LARGE CHAIN (EC 6.3.5.5)	CYTOSINE DEAMINASE (EC 3.5.4.1)	CYTOSINE DEAMINASE (EC 3.5.4.1)	CYTOSINE DEAMINASE (EC 3 5.4.1)	CREATININE DEAMINASE (EC 3.5.4.21)	DEOXYCYTIDINE TRIPHOSPHATE DEAMINASE (EC 3.5.4.13)	THYMIDYLATE SYNTHASE (EC 2.1.1.45)	URACIL PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.9)	URACIL PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.9)
NT Stop	10900	8193	9589	1003	1142	4040	3748	775	17346	7013	3362	5283	10441	28046	3198	34814	2	16810	7935	2341	9579	1080	1082
NT Start	9722	7258	8249	2	591	3207	3020	47	16672	7621	3769	4576	8780	24708	-	34491	322	15566	6691	1862	9680	568	920
Contig.	GR00022	GR00022	GR00022	GR00647	GR00462	GR00654	VV0150	GR00542	GR00014	GR00020	GR00040	GR00188	GR00447	VV0134	GR00654	W0112	GR00110	VV 0020	GR00655	W0237	W0129	VV0328	GR10003
Identification Code	RXA00147	RXA00145	RXA00146	RXA02208	RXA01660	RXA02235	RXN01892	F RXA01892	RXA00105	RXA00131	RXA00266	RXA00718	RXA01599	RXN02234	F RXA02234	RXN00450	F RXA00450	RXN02272	F RXA02272	RXN03004	RXN03137	RXN03171	F RXA02857
Amino Acid SEQ ID NO	974	976	978	980	982	984	986	988	066	992	964	966	866	1000	1002	1004	1006	1008	1010	1012	1014	1016	1018
Nucleic Acid	973	975	226	676	981	983	985	286	686	991	993	995	266	666	1001	1003	1005	1007	1009	1011	1013	1015	1017

Purine and pyrimidine base, nucleoside and nucleotide salvage, interconversion, reduction and degradation: Table 1 (continued) Purines:

Function A PARAMETER A CE VEC 2 4 2 3 3	ADENINE PHOSPHORIBOSTLI RANSFERASE (EC. 2.4 2.7) HYPOXANTHINE-GUANINE PHOSPHORIBOSYLTRANSFERASE (EC. 2.4.2.8)	XANTHINE-GUANINE PHOSPHORIBOSYLTRANSFERASE (EC 2.4.2.22)	GTP PYROPHOSPHOKINASE (EC 2.7.6.5)	GUANOSINE-3',5'-BIS(DIPHOSPHATE) 3'-PYROPHOSPHOHYDROLASE (EC	3.1.7.2) GUANOSINE-3',5'-BIS(DIPHOSPHATE) 3'-PYROPHOSPHOHYDROLASE (EC	3.1.7.2) GLIANOSINE-3: 5-BIS(DIPHOSPHATE) 3:-PYROPHOSPHOHYDROLASE (EC	3.17.2) GUANOSINE-3:5-BIS(DIPHOSPHATE) 3: PYROPHOSPHOHYDROLASE (EC	3.1.7.2) DEOXYGUANOSINETRIPHOSPHATE TRIPHOSPHOHYDROLASE (EC.3.1.5.1)	DIADENOSINE 5',5"-P1,P4-TETRAPHOSPHATE HYDROLASE (EC 3.6.1.17)	DIADENOSINE 5',5"-P1,P4-TETRAPHOSPHATE HYDROLASE (EC 3.6.1.17)	DIADENOSINE 5',5"-P1,P4-TETRAPHOSPHATE HYDROLASE (EC 3.6.1.17)	DIADENOSINE 5',5""-P1,P4-TETRAPHOSPHATE HYDROLASE (EC 3.6.1.17)	PHOSPHOADENOSINE PHOSPHOSULFATE REDUCTASE (EC 1.8.99.4)	DIMETHYLADENOSINE TRANSFERASE (EC 2.1.1)	AMP NUCLEOSIDASE (EC 3.2.2.4)	AMP NUCLEOSIDASE (EC 3.2.2.4)	GTP PYROPHOSPHOKINASE (EC 2.7.6.5)	GUANOSINE-3',5'-BIS(DIPHOSPHATE) 3'-PYROPHOSPHOHYDROLASE (EC	3.1.7.2)
NT Stop	1883	3347	4017	1011	2741	2802	3677	18240	6768	5	2347	5126	9	2117	3323	34	29420	2	
NT Start	1329	3820	3388	2045	1962	2741	3147	19511	5761	661	2580	5653	446	1239	1893	1101	30442	1138	
Contig	GR00772 GR00424	GR00618	GR00276	VV0171	GR00772	GR00772	GR00517	GR00422	VV0143	GR00293	GR00294	GR00425	GR00012	GR00537	VV0152	GR00659	0600//	VV0171	
Identification Code	RXA62771 RXA01512	RXA02031	RXA00981	RXN02772	F RXA02772	F RXA02773	RXA01835	EXA01483	RXN01027	F RXA01024	F RXA01027	RXA01528	RXA00072	RXA01878	RXN02281	F RXA02281	RXN01240	RXN02008	
Amino Acid SEQ ID NO	1020	1024	1026	1028	1030	1032	1034	1036	1038	1040	1042	1044	1046	1048	1050	1052	1054	1056	
D AC	1019					1031	1031	1035	1037	1039	1041	1043	1045	1047	1049	1051	1053	1055	

Pyrimdine and purine metabolism:

Function	INOSINE-URIDINE PREFERRING NUCLEOSIDE HYDROLASE (EC 3.2.2.1)	INOSINE-URIDINE PREFERRING NUCLEOSIDE HYDROLASE (EC 3.2.2.1)	INOSINE-URIDINE PREFERRING NUCLEOSIDE HYDROLASE (EC 3.2.2.1)	EXOPOLYPHOSPHATASE (EC 3.6.1.11)	RIBONUCLEOSIDE-DIPHOSPHATE REDUCTASE ALPHA CHAIN (EC 1.17.4.1)	RIBONUCLEOSIDE-DIPHOSPHATE REDUCTASE ALPHA CHAIN (EC 1.17.4.1)	RIBONUCLEOSIDE-DIPHOSPHATE REDUCTASE ALPHA CHAIN (EC 1.17.4.1)	RIBONUCLEOSIDE-DIPHOSPHATE REDUCTASE 2 BETA CHAIN (EC 1.17.4.1)	RIBONUCLEOTIDE REDUCTASE SUBUNIT R2F	NRDI PROTEIN	POLYRIBONUCLEOTIDE NUCLEOTIDYLTRANSFERASE (EC 2.7.7.8)	POLYRIBONUCLEOTIDE NUCLEOTIDYLTRANSFERASE (EC 2.7.7.8)	POLYRIBONUCLEOTIDE NUCLEOTIDYLTRANSFERASE (EC 2.7.7.8)
NT Stop	9333	581	6320	10985	35982	4	2062	31842	806	797	627	631	4
NT Start	10268	ဗ	5418	10059	38084	693	3402	32843	1321	1240	-	2	099
Contig	VV0120	GR00557	GR00731	GR00720	VV0084	GR00301	GR00302	VV0084	GR00550	GR00301	GR00237	GR00413	GR00423
Identification Code	RXN01940	F RXA01940	RXA02559	RXA02497	RXN01079	F RXA01079	F RXA01084	RXN01920	F RXA01920	RXA01080	RXA00867	RXA01416	RXA01486
Amino Acid	1058	1060	1062	1064	1066	1068	1070	1072	1074	1076	1078	1080	1082
Nucleic Acid	1057	1059	1061	1063	1065	1067	1069	1071	1073	1075	1077	1079	1081

	2'3'-CYCLIC-NUCLEOTIDE 2'-PHOSPHODIESTERASE (EC 3.1.4.16) 2'3'-CYCLIC-NUCLEOTIDE 2'-PHOSPHODIESTERASE (EC 3.1.4.16) 2'3'-CYCLIC-NUCLEOTIDE 2'-PHOSPHODIESTERASE (EC 3.1.4.16) INOSINE-URIDINE PREFERRING NUCLEOSIDE HYDROLASE (EC 3.2.2.1) CYTOSOLIC PROTEIN INVOLVED IN PURINE METABOLISM PROTEIN INVOLVED IN METABOLISM PROTEIN INVOLVED IN METABOLISM PROTEIN INVOLVED IN METABOLISM OF S-ADENOSYLMETHIONINE, PURINES AND PANTOTHENATE ABC TRANSPORTER ATP-BINDING PROTEIN INVOLVED IN PURINE METABOLISM	Function URACIL PHOSPHORIBOSYLTRANSFERASE (EC 2 4 2 9) CYTOSINE DEAMINASE (EC 3 5 4 1) CYTOSONE DEAMINASE (EC 3 5 4 1) RIBOSOMAL LARGE SUBUNIT PSEUDOURIDINE SYNTHASE B (EC 4 2 1 7 0) PHOSPHATIDATE CYTIDYLYLTRANSFERASE (EC 2 7 7 4 7) PHOSPHOMETHYLPYRIMIDINE KINASE (EC 2 7 4 7) PHOSPHOMETHYLPYRIMIDINE KINASE (EC 2 7 4 7) PHOSPHOMETHYLPYRIMIDINE KINASE (EC 2 7 4 7) CYTOSOLIC PROTEIN INVOLVED IN PYRIMIDINE METABOLISM CYTOSOLIC PROTEIN INVOLVED IN METABOLISM OF PYRIDIMES AND ADENOSYLHOMOCYSTEINE CYTOSOLIC PROTEIN INVOLVED IN PYRIMIDINE METABOLISM EXPORTED PROTEIN INVOLVED IN PYRIMIDINE METABOLISM CYTOSOLIC PROTEIN INVOLVED IN PYRIMIDINE METABOLISM
Function	2;3-CYCLIC-N 2;3-CYCLIC-N INOSINE-URII CYTOSOLIC F PROTEIN INV CYTOSOLIC F MEMBRANE S PROTEIN INV LIPOPROTEIN INV PROTEIN INV AND PANTOT ABC TRANSP METABOLISM	Eunction URACIL URACIL CYTOSII CYTOSII CYTOSII CYTOSI RIBOSO PHOSPP PHOSPP PHOSPP PHOSPP PHOSPP PHOSPP CYTOSC
Table 1 (continued)	7689 8964 40789	NT Stop 1080 1082 34814 5 828 4576 2476 2446 2246 22858 616
Ta NT Start	7162 7729 39842	NT Start 568 570 34491 322 337 3617 1622 8581 1019 22187 2
Contig	GR00467 GR00467 VV0139	Contig. VV0328 GR10003 VV0112 GR00110 GR0018 GR00542 GR00542 GR00726 VV0270 GR00348
Identification Code	RXA01678 RXA01679 RXN01488 RXC00540 RXC00560 RXC01088 RXC02624 RXC02665 RXC02238 RXC02238	Identification Code RXN03171 F RXA02857 RXN00450 F RXA00450 F RXA00450 RXA00450 RXA00177 RXA01894 RXA01209 F RXA01209 F RXA01209 F RXA01209 F RXA01209 RXC01600 RXC01600 RXC01600 RXC01709 RXC01709 RXC01709
Amino Acid	1084 1086 1086 1090 1092 1096 1100 1100 1100 1100 1100	Amino Acid SEO ID NO 1106 1110 1112 1112 1124 1126 1127 1136 1136 1136 1136
Nucleic Acid	1083 1083 1083 1088 1088 1088 1088 1089 1091 1101 11101 11103 1103 1103 1103 1103 1103 1103 1103 1103 1103 1103 1103 110	Nucleic Acid SEQ ID NO 1105 1105 1110 1111 1111 1112 1126 1127 1131 1133 1135

Table 1 (continued)

Sugars Trehalose

Function TREHALOSE-PHOSPHATASE (EC 3.1.3.12) maltooligosyltrehalose synthase maltooligosyltrehalose synthase maltooligosyltrehalose trehalose terbalotrolase TREHALOSE/MALTOSE BINDING PROTEIN Hypothetical Trehalose-Binding Protein Hypothetical Trehalose Transport Protein TREHALOSE/MALTOSE BINDING PROTEIN TREHALOSE/MALTOSE BINDING PROTEIN TREHALOSE/MALTOSE BINDING PROTEIN TRANSMEBRANE PROTEIN INVOLVED IN TREHALOSE METABOLISM
NT Stop 1013 30489 7579 2543 4 39017
NT Start 246 32921 5147 714 735 38532
Contig. GR00065 VV0090 GR00358 GR00751 VV0051
Identification Code RXA00347 RXN01239 F RXA01239 F RXA02845 RXN02355 RXN02355 RXN02369 RXS00349 RXS003183 RXC00874
Amino Acid SEQ ID NO 1140 1144 1146 1148 1150 1152 1154
Nucleic Acid SEQ ID NO 1139 1141 1145 1147 1149 1151

			TABLE 2 - Excluded Genes	ded Genes
GenBank™	Gene Name	Gene Function	Į.	Reference
Accession No.				
A09073	Яdd	Phosphoerol p	ol pyruvate carboxylase	Bachmann, B. et al. "DNA fragment coding for phosphoenolpyruvat corboxylase, recombinant DNA carrying said fragment, strains carrying the recombinant DNA and method for producing L-aminino acids using said strains," Patent: EP 0358940-A 3 03/21/90
A45579, A45581, A45583, A45585		Threonine Jeh	lehydratase	Moeckel, B. et al. "Production of L-isoleucine by means of recombinant micro-organisms with deregulated threonine dehydratase," Patent: WO 9519442-A 5 07/20/95
AB003132	murC; ftsQ; ftsZ			Kobayashi, M. et al. "Cloning, sequencing, and characterization of the fts7. gene from coryneform bacteria," <i>Biochem. Biophys. Res. Commun.</i> , 236(2):383-388 (1997)
AB015023	murC; ftsQ			Wachi, M. et al. "A murC gene from Coryneform bacteria," Appl. Microbiol. Biotechnol., 51(2):223-228 (1999)
AB018530	dtsR			Kimura, E. et al. "Molecular cloning of a novel gene, dtsR, which rescues the detergent sensitivity of a mutant derived from <i>Brevibacterium</i> lactofermentum," Biosci. Biotechnol ochem., 60(10):1565-1570 (1996)
AB018531	dtsR1; dtsR2			
AB020624	murl	D-glutamate ra	racemase	
AB023377	tkt	transketola		
AB024708	gltB; gltD	Glutamine -o.	-oxoglutarate aminotransferase	
AB025424	acn	aconitase		
AB027714	rep	Replication pro	protein	
AB027715	rep; aad	_	protein; aminoglycoside erase	
AF005242	argC	N-acetylglu am dehydrogen se	amate-5-semialdehyde Ise	
AF005635	glnA	-	ynthetase	
AF030405	hisF	cyclase		
AF030520	argG	Argininosucin	cinate synthetase	
AF031518	argF	-	rbamolytransferase	
AF036932	aroD	3-dehydroq in	inate dehydratase	
AF038548	pyc	Pyruvate ca bo	boxylase	

		Table 2 (continued)	(penu
AF038651	dciAE; apt; rel	Dipeptide-binding protein; adenine phosphoribosyltransferase; GTP pyrophosphokinase	Wehmeier, L. et al. "The role of the Corynebacterium glutamicum rel gene in (p)ppGpp metabolism," <i>Microbiology</i> , 144:1853-1862 (1998)
AF041436	argR	Arginine repressor	
AF045998	Admi	Inositol monophosphate phosphatase	
AF048764	areH	Argininosuccinate lyase	
AF049897	argC; argJ; argB;	N-acetylglutamylphosphate reductase;	
	argU; argr; argr;	acetylglutamate kinase, acetylomithine	
	; ; ; ; ; ; ; ; ; ; ; ; ; ; ; ; ; ; ;	transminase; ornithine	
		carbamoyltransferase; arginine repressor;	
		argininosuccinate synthase;	
		argininosuccinate lyase	
AF050109	inhA	Enoyl-acyl carrier protein reductase	
AF050166	hisG	ATP phosphoribosyltransferase	
AF051846	hisA	Phosphoribosylformimino-5-amino-1-	
		phosphoribosyl-4-imidazolecarboxamide	
		tagnicias.	Date of all steplation and analysis of met A a methionine biosynthetic gene
AF052652	metA	Homoserine O-acetyltransferase	Fark, S. et al. Isolation and analysis of moon, a mooning the encoding homoserine acetyltransferase in Corynebacterium glutamicum," Mol. Cells, 8(3):286-294 (1998)
AF053071	aroB	Dehydroquinate synthetase	
A F060558	hisH	Glutamine amidotransferase	
AF086704	hisE	Phosphoribosyl-ATP-	
		pyrophosphonyurorase	
AF114233	aroA	5-enolpyruvylshikimate 3-phosphate synthase	Olivia Band Dunina
AF116184	panD	L-aspartate-alpha-decarboxylase precursor	Dusch, N. et al. "Expression of the Corynebacterium glutamicum pair Dusch, N. et al. "Expression of the Corynebacterium glutamicum pair Denie encoding L-aspartate-alpha-decarboxylase leads to pantothenate overproduction in Escherichia coli," <i>Appl. Environ. Microbiol.</i> , 65(4)1530-1539 (1999)
AF124518	aroD; aroE	3-dehydroquinase; shikimate dehydrogenase	
AF124600	aroC; aroK; aroB; pepQ	Chorismate synthase; shikimate kinase; 3-dehydroquinate synthase; putative cytoplasmic peptidase	
AF145897	inhA		
A E 1 4 5 0 0 9	Adai		
AF143696	Cilli		

			Table 2 (continued)	(Political)
AJ001436	ectP	Transport proline	f ectoine, glycine betaine,	Peter, H. et al. "Corynebacterium glutamicum is equipped with four secondary carriers for compatible solutes: Identification, sequencing, and characterization of the proline/ectoine uptake system, ProP, and the ectoine/proline/glycine betaine carrier, EctP," J. Bacteriol, 180(22):6005-6012 (1998)
AJ004934	дарД	Tetrahydro (incomplet	dipicolinate succinylase	Wehrmann, A. et al. "Different modes of diaminopimelate synthesis and their role in cell wall integrity: A study with Corynebacterium glutamicum," J. Bacteriol., 180(12):3159-3165 (1998)
AJ007732	ppc; secG; amt; ocd; soxA	Phosphoen affinity am ornithine-c oxidase	olpyruvate-carboxylase; ?; high monium uptake protein; putative rclodecarboxylase; sarcosine	
AJ010319	ftsY, glnB, glnD, srp; amtP	Involved ir uridylyltrat enzmye); s affinity am	cell division; PII protein; sferase (uridylyl-removing gnal recognition particle; low nonium uptake protein	Jakoby, M. et al. "Nitrogen regulation in Corynebacterium glutamicum; Isolation of genes involved in biochemical characterization of corresponding proteins," FEMS Microbiol, 173(2):303-310 (1999)
AJ132968	cat	Chloramph	enicol aceteyl transferase	
AJ224946	овш	L-malate: 0	quinone oxidoreductase	Molenaar, D. et al. "Biochemical and genetic characterization of the membrane-associated malate dehydrogenase (acceptor) from Corynebacterium glutamicum," Eur. J. Biochem., 254(2):395-403 (1998)
AJ238250	ndh	NADH deh	ydrogenase	
AJ238703	porA	Porin		Lichtinger, T. et al. "Biochemical and biophysical characterization of the cell wall porin of Corynebacterium glutamicum: The channel is formed by a low molecular mass polypeptide," <i>Biochemistry</i> , 37(43):15024-15032 (1998)
D17429		Transposab	e element 1S31831	Vertes et al. "Isolation and characterization of IS31831, a transposable element from Corynebacterium glutamicum," Mol. Microbiol., 11(4):739-746 (1994)
D84102	odhA	2-oxogluta	rate dehydrogenase	Usuda, Y. et al. "Molecular cloning of the Corynebacterium glutamicum (Brevibacterium lactofermentum AJ12036) odhA gene encoding a novel type of 2-oxoglutarate dehydrogenase," <i>Microbiology</i> , 142.3347-3354 (1996)
E01358	hdh; hk	Homoserin kinase	dehydrogenase, homoserine	Katsumata, R. et al. "Production of L-thereonine and L-isoleucine," Patent: JP 1987232392-A 1 10/12/87
E01359		Upstream o	of the start codon of homoserine	Katsumata, R. et al. "Production of L-thereonine and L-isoleucine," Patent: JP 1987232392-A 2 10/12/87
E01375		Tryptophan	n operon	
E01376	trpL; trpE	Leader pep	de; anthranilate synthase	Matsui, K. et al. "Tryptophan operon, peptide and protein coded thereby, utilization of tryptophan operon gene expression and production of tryptophan," Patent. JP 1987244382-A 1 10/24/87

	Table 2 (continued)	nued)
E01377	Promoter and operator regions of tryptophan operon	Matsui, K. et al. "Tryptophan operon, peptide and protein coded thereby, utilization of tryptophan operon gene expression and production of tryptophan," Patent: JP 1987244382-A 1 10/24/87
E03937	Biotin-synthase	Hatakeyama, K. et al. "DNA fragment containing gene capable of coding biotin synthetase and its utilization," Patent: JP 1992278088-A 1 10/02/92
E04040	Diamino pelargonic acid aminotransferase	Kohama, K. et al. "Gene coding diaminopelargonic acid aminotransferase and desthiobiotin synthetase and its utilization," Patent: JP 1992330284-A 1 11/18/92
E04041	Desthiobiotinsynthetase	Kohama, K. et al. "Gene coding diaminopelargonic acid aminotransferase and desthiobiotin synthetase and its utilization," Patent: JP 1992330284-A 1 11/18/92
E04307	Flavum aspartase	Kurusu, Y. et al. "Gene DNA coding aspartase and utilization thereof," Patent: JP 1993030977-A 1 02/09/93
E04376	Isocitric acid lyase	Katsumata, R. et al. "Gene manifestation controlling DNA," Patent: JP 1993056782-A 3 03/09/93
E04377	Isocitric acid lyase N-terminal fragment	Katsumata, R. et al. "Gene manifestation controlling DNA," Patent: JP 1993056782-A 3 03/09/93
E04484	Prephenate dehydratase	Sotouchi, N. et al. "Production of L-phenylalanine by fermentation," Patent: JP 1993076352-A 2 03/30/93
E05108	Aspartokinase	Fugono, N. et al. "Gene DNA coding Aspartokinase and its use," Patent: JP 1993184366-A 1 07/27/93
E05112	Dihydro-dipichorinate synthetase	Hatakeyama, K. et al. "Gene DNA coding dihydrodipicolinic acid synthetase and its use," Patent: JP 1993184371-A 1 07/27/93
E05776	Diaminopimelic acid dehydrogenase	Kobayashi, M. et al. "Gene DNA coding Diaminopimelic acid dehydrogenase and its use," Patent: JP 1993284970-A 1 11/02/93
E05779	Threonine synthase	Kohama, K. et al. "Gene DNA coding threonine synthase and its use," Patent: JP 1993284972-A 1 11/02/93
E06110	Prephenate dehydratase	Kikuchi, T. et al. "Production of L-phenylalanine by fermentation method," Patent: JP 1993344881-A 1 12/27/93
E06111	Mutated Prephenate dehydratase	Kikuchi, T. et al. "Production of L-phenylalanine by fermentation method," Patent: JP 1993344881-A 1 12/27/93
E06146	Acetohydroxy acid synthetase	Inui, M. et al. "Gene capable of coding Acetohydroxy acid synthetase and its use," Patent: JP 1993344893-A 1 12/27/93
E06825	Aspartokinase	Sugimoto, M. et al. "Mutant aspartokinase gene," patent: JP 1994062866-A 1 03/08/94
E06826	Mutated aspartokinase alpha subunit	Sugimoto, M. et al. "Mutant aspartokinase gene," patent: JP 1994062866-A 1 03/08/94

		Table 2 (continued)	linued)
E06827		Mutated a partokinase alpha subunit	Sugimoto, M. et al. "Mutant aspartokinase gene," patent. JP 1994062866-A 1 03/08/94
E07701	secY		Honno, N. et al. "Gene DNA participating in integration of membraneous protein to membrane," Patent: JP 1994169780-A 1 06/21/94
E08177	As	Aspartokirase	Sato, Y. et al. "Genetic DNA capable of coding Aspartokinase released from feedback inhibition and its utilization," Patent: JP 1994261766-A 1 09/20/94
E08178, E08179, E08180, E08181, E08182	Fe	Feedback hhibition-released Aspartokinase	Sato, Y. et al. "Genetic DNA capable of coding Aspartokinase released from feedback inhibition and its utilization," Patent: JP 1994261766-A 1 09/20/94
E08232	Ac	Acetohydroxy-acid isomeroreductase	Inui, M. et al. "Gene DNA coding acetohydroxy acid isomeroreductase," Patent: JP 1994277067-A 1 10/04/94
E08234	Esec		Asai, Y. et al. "Gene DNA coding for translocation machinery of protein," Patent: JP 1994277073-A 1 10/04/94
E08643	FT	FT aminotransferase and desthiobiotin synthetase promoter region	Hatakeyama, K. et al. "DNA fragment having promoter function in coryneform bacterium," Patent: JP 1995031476-A 1 02/03/95
E08646	Bi	Biotin synthetase	Hatakeyama, K. et al. "DNA fragment having promoter function in coryneform bacterium," Patent: JP 1995031476-A 1 02/03/95
E08649	As	Aspartase	Kohama, K et al "DNA fragment having promoter function in coryneform bacterium," Patent: JP 1995031478-A 1 02/03/95
E08900	.iQ	Dihydrodipicolinate reductase	Madori, M. et al. "DNA fragment containing gene coding Dihydrodipicolinate acid reductase and utilization thereof," Patent: JP 1995075578-A 1 03/20/95
E08901	Di	Diaminopinelic acid decarboxylase	Madori, M. et al. "DNA fragment containing gene coding Diaminopimelic acid decarboxylase and utilization thereof," Patent: JP 1995075579-A 1 03/20/95
E12594	Se	Serine hydoxymethyltransferase	Hatakeyama, K. et al. "Production of L-trypophan," Patent: JP 1997028391-A 1 02/04/97
E12760, E12759, E12758	tra	transposase	Moriya, M. et al. "Amplification of gene using artificial transposon," Patent: JP 1997070291-A 03/18/97
E12764	Ar	Arginyl-tR NA synthetase; diaminopimelic acid decarboxylase	Moriya, M. et al. "Amplification of gene using artificial transposon," Patent: JP 1997070291-A 03/18/97
E12767	IQ	Dihydrodip colinic acid synthetase	Moriya, M. et al. "Amplification of gene using artificial transposon," Patent: JP 1997070291-A 03/18/97
E12770	asi	aspartokinake	Moriya, M. et al. "Amplification of gene using artificial transposon," Patent: JP 1997070291-A 03/18/97
E12773	O	Dihydrodip colinic acid reductase	Moriya, M. et al. "Amplification of gene using artificial transposon," Patent: JP 1997070291-A 03/18/97

		Table 2 (continued)	(penu
E13655		Glucose-6-phosphate dehydrogenase	Hatakeyama, K. et al. "Glucose-6-phosphate dehydrogenase and DNA capable of coding the same," Patent: JP 1997224661-A 1 09/02/97
L01508	llvA	Threonine dehydratase	Moeckel, B. et al. "Functional and structural analysis of the threonine dehydratase of Corynebacterium glutamicum," <i>J. Bacteriol.</i> , 174:8065-8072 (1992)
1.07603	EC 4.2.1.15	3-deoxy-D-arabinoheptulosonate-7- phosphate synthase	Chen, C. et al. "The cloning and nucleotide sequence of Corynebacterium glutamicum 3-deoxy-D-arabinoheptulosonate-7-phosphate synthase gene," <i>FEMS Microbiol. Lett.</i> , 107:223-230 (1993)
1.09232	IIvB; iIvN; iIvC	Acetohydroxy acid synthase large subunit; Acetohydroxy acid synthase small subunit; Acetohydroxy acid isomeroreductase	Keilhauer, C. et al. "Isoleucine synthesis in Corynebacterium glutamicum. molecular analysis of the ilvB-ilvN-ilvC operon," <i>J. Bacteriol.</i> , 175(17):5595-5603 (1993)
1.18874	PtsM	Phosphoenolpyruvate sugar phosphotransferase	Fouet, A et al. "Bacillus subtilis sucrose-specific enzyme II of the phosphotransferase system: expression in Escherichia coli and homology to enzymes II from enteric bacteria," <i>PNAS USA</i> . 84(24):8773-8777 (1987); Lee, J.K. et al. "Nucleotide sequence of the gene encoding the Corynebacterium glutamicum mannose enzyme II and analyses of the deduced protein sequence," <i>FEMS Microbiol. Lett.</i> , 119(1-2):137-145 (1994)
L27123	aceB	Malate synthase	Lee, H-S. et al. "Molecular characterization of aceb, a gene encoding matate synthase in Corynebacterium glutamicum," J. Microbiol. Biotechnol., 4(4):256-263 (1994)
1.27126		Pyruvate kinase	Jetten, M. S. et al. "Structural and functional alialysis of pyruvaic reflections." Appl. Environ. Microbiol., 60(7):2501-2507 (1994)
L28760	aceA	Isocitrate lyase	o the Annual Control DNA common analysis and
L35906	dtxr	Diphtheria toxin repressor	Oguiza, J.A. et al. "Molecular cioning, Diva sequence anarysis, and characterization of the Corynebacterium diphtheriae dtxR from Brevibacterium lactofermentum," <i>J. Bacteriol.</i> , 177(2):465-467-467 (1995)
M13774		Prephenate dehydratase	Follettie, M.T. et al. "Molecular cioning and nucleotide sequence of the Corynebacterium glutamicum pheA gene," J. Bacteriol., 167:695-702 (1986)
M16175	5S rRNA		Park, Y-H. et al. "Phylogenetic alialysis of the colyncial caceria of colyncial rRNA sequences," J. Bacteriol., 169:1801-1806 (1987)
M16663	trpE	Anthranilate synthase, 5' end	Sano, K. et al. "Structure and function of the up operon control regions of Brevibacterium lactofermentum, a glutamic-acid-producing bacterium," Gene. 52:191-200 (1987)
M16664	trpA	Tryptophan synthase, 3'end	Sano, K. et al. "Structure and function of the tip operous of Brevibacterium lactofermentum, a glutamic-acid-producing bacterium," Gene. 52:191-200 (1987)

			Table 2 (continued)	(panu
M25819		Phosphoen	olpyruvate carboxylase	O'Regan, M. et al. "Cloning and nucleotide sequence of the
				Phosphoenolpyruvate carboxylase-coding gene of Corynebacterium glutamicum ATCC13032." Gene. 77(2):237-251 (1989)
M85106		23S rRNA	gene insertion sequence	Roller, C. et al. "Gram-positive bacteria with a high DNA G:C content are characterized by a common insertion within their 23S rRNA genes," J. Gen.
		_		Microbiol., 138:1167-1175 (1992)
M85107,		23S rRNA	gene insertion sequence	Roller, C. et al. "Gram-positive bacteria with a high DNA G+C content are
M85108				characterized by a common insertion within their 23S rRNA genes," J. Gen. Microbiol., 138:1167-1175 (1992)
M89931	aecD; brnQ; yhbw	Beta C-S ly	ase; branched-chain amino acid	Rossol, I. et al. "The Corynebacterium glutamicum accD gene encodes a C-S
		uptake carr	er; hypothetical protein yhbw	lyase with alpha, beta-elimination activity that degrades aminoethyleysteine,"
				J. Bacteriol., 174(9):2968-2977 (1992); Tauch, A. et al. "Isoleucine uptake in
				Corynebacterium glutamicum ATCC 13032 is directed by the brnQ gene product," Arch Microbiol, 169(4):303-312 (1998)
S59299	tra	Leader gen	(promoter)	Herry, D.M. et al. "Cloning of the trp gene cluster from a tryptophan-
	•)		hyperproducing strain of Corynebacterium glutamicum: identification of a
				mutation in the trp leader sequence," Appl Environ Microbiol, 59(3):791-799
				(1993)
U11545	трД	Anthranilat	phosphoribosyltransferase	O'Gara, J.P. and Dunican, L.K. (1994) Complete nucleotide sequence of the
				Coryncoacterium gintamicum A I C. 2 1830 4pD gene. Thesis, Microbiology Department University College Galway Ireland
U13922	cellM; cellR; clellR	Putative type	e II 5-cytosoine	Schafer, A. et al. "Cloning and characterization of a DNA region encoding a
)	methyltrans		stress-sensitive restriction system from Corynebacterium glutamicum ATCC
		restriction		13032 and analysis of its role in intergeneric conjugation with Escherichia
		type III rest		coli," J. Bacteriol., 176(23):7309-7319 (1994); Schafer, A. et al. "The
				Corynebacterium glutamicum cgIIM gene encoding a 5-cytosine in an McrBC-deficient Escherichia coli strain," <i>Gene</i> , 203(2):95-101 (1997)
U14965	recA			
U31224	xdd			Ankri, S. et al. "Mutations in the Corynebacterium glutamicumproline biosynthetic pathway: A natural bypass of the proA step," J. Bacteriol., 178(15):4412,4419 (1996).
U31225	proC	L-proline:	ADP+ 5-oxidoreductase	Ankri, S. et al. "Mutations in the Corynebacterium glutamicumproline
		-		biosynthetic pathway: A natural bypass of the proA step," J. Bacteriol., 178(15) 4412-4419 (1996)
U31230	obg; proB; unkdh	?;gamma g	stamyl kinase;similar to D.	Ankri, S. et al. "Mutations in the Corynebacterium glutamicumproline
	•	isomer spec		biosynthetic pathway: A natural bypass of the proA step," J. Bacteriol., 178(15):4412-4419 (1996)
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(pan	Serebriiskii, I.G., "Two new members of the bio B superfamily: Clohing, sequencing and expression of bio B genes of Methylobacillus flagellatum and Corynebacterium glutamicum," <i>Gene.</i> 175:15-22 (1996)	Jager, W. et al. "A Corynebacterium glutamicum gene encoding a two-domain protein similar to biotin carboxylases and biotin-carboxyl-carrier proteins," <i>Arch. Microbiol.</i> , 166(2):76-82 (1996)	Jager, W. et al. "A Corynebacterium glutamicum gene conferring multidrug resistance in the heterologous host Escherichia coli," J. Bacteriol., 179(7):2449-2451 (1997)				Matsui, K. et al. "Complete nucleotide and deduced amino acid sequences of the Brevibacterium lactofermentum tryptophan operon," <i>Nucleic Acids Res.</i> 14(24):10113-10114 (1986)	Yeh, P. et al. "Nucleic sequence of the lysA gene of Corynebacterium glutamicum and possible mechanisms for modulation of its expression," Mol. Gen. Genet., 212(1):112-119 (1988)	Eikmanns, B.J. et al. "The Phosphoenolpyruvate carboxylase gene of Corynebacterium glutamicum: Molecular cloning, nucleotide sequence, and expression," <i>Mol. Gen. Genet.</i> , 218(2):330-339 (1989); Lepiniec, L. et al. "Sorghum Phosphoenolpyruvate carboxylase gene family: structure, function and molecular evolution," <i>Plant. Mol. Biol.</i> , 21 (3):487-502 (1993)	Von der Osten, C.H. et al. "Molecular cloning, nucleotide sequence and fine- structural analysis of the Corynebacterium glutamicum fda gene: structural comparison of C. glutamicum fructose-1, 6-biphosphate aldolase to class I and class II aldolases," Mol. Microbiol.	Bonnassie, S. et al. "Nuclete sequence of the dapA gene from Corynebacterium glutamicum," Nucleic Acids Res., 18(21):6421 (1990)	Cianciotto, N. et al. "DNA sequence homology between att B-related sites of Corynebacterium diphtheriae, Corynebacterium ulcerans, Corynebacterium glutamicum, and the attP site of lambdacorynephage," FEMS. Microbiol, Lett., 66:299-302 (1990)	Marcel, T. et al. "Nucleotide sequence and organization of the upstream region of the Corynebacterium glutamicum lysA gene," <i>Mol. Microbiol.</i> , 4(11):1819-1830 (1990)
Table 2 (continued)	Biotin synthase	Thiosulfate sulfurtransferase; acyl CoA carboxylase	Multidrug resistance protein	Heat shock ATP-binding protein	3'5"-aminoglycoside phosphotransferase	Corynebacterium glutamicum unidentified sequence involved in histidine biosynthesis, partial sequence	Tryptophan operon	DAP decarboxylase (meso-diaminopimelate decarboxylase, EC 4.1.1.20)	Phosphoenolpyruvate carboxylase	Fructose-bisphosphate aldolase	L-2, 3-dihydrodipicolinate synthetase (EC 4.2.1.52)	l ate	Arginyl-tRNA synthetase; Diaminopimelate decarboxylase
	bioB	thtR; accBC	cmr	clpB	aphA-3		trpA; trpB; trpC; trpD; trpE; trpG; trpL	lys A	EC 4.1.1.31	fda	dapA		argS; lysA
	U31281	U35023	U43535	U43536	U53587	U89648	X04960	X07563	X14234	X17313	X53993	X54223	X54740

			Table 2 (continued)	(poned)
X55994	trpL, trpE	Putative le	e; anth	Heery, D.M. et al. "Nucleotide sequence of the Corynebacterium glutamicum
		synthase c	omponent l	trpl: gene," Nucleic Acids Res., 18(23):7138 (1990)
X56037	thrC	Threonine	synthase	Han, K.S. et al. "The molecular structure of the Corynebacterium glutamicum threonine synthase gene," Mol. Microbiol., 4(10):1693-1702 (1990)
X56075	attB-related site	Attachmer	t site	Cianciotto, N. et al. "DNA sequence homology between att B-related sites of Corynebacterium diphtheriae, Corynebacterium ulcerans, Corynebacterium glutamicum, and the attP site of lambdacorynephage," FEMS. Microbiol, Lett., 66:299-302 (1990)
X57226	lysC-alpha; lysC-beta; asd	Aspartokir Aspartokir semialdeh	ase-alpha subunit; ase-beta subunit; aspartate beta de dehydrogenase	Kalinowski, J. et al. "Genetic and biochemical analysis of the Aspartokinase from Corynebacterium glutamicum," <i>Mol. Microbiol</i> , 5(5):1197-1204 (1991); Kalinowski, J. et al. "Aspartokinase genes lysC alpha and lysC beta overlap and are adjacent to the aspertate beta-semialdehyde dehydrogenase gene asd in Corynebacterium glutamicum," <i>Mol. Gen. Genet.</i> , 224(3):317-324 (1990)
X59403	gap;pgk; tpi	Glyceralde phosphogl isomerase	nyde-3-phosphate; cerate kinase; triosephosphate	Eikmanns, B.J. "Identification, sequence analysis, and expression of a Corynebacterium glutamicum gene cluster encoding the three glycolytic enzymes glyceraldehyde-3-phosphate dehydrogenase, 3-phosphoglycerate kinase, and triosephosphate isomeras," J. Bacteriol, 174(19):6076-6086 (1992)
X59404	gdh	Glutamate	lehydrogenase	Bormann, E.R. et al. "Molecular analysis of the Corynebacterium glutamicum gdh gene encoding glutamate dehydrogenase," Mol. Microbiol., 6(3):317-326 (1992)
X60312	lysI	L-lysine po	rmease	Seep-Feldhaus, A.H. et al. "Molecular analysis of the Corynebacterium glutamicum lysl gene involved in lysine uptake," Mol. Microbiol., 5(12):2995-3005 (1991)
X66078	cop1	Ps1 protein		Joliff, G. et al. "Cloning and nucleotide sequence of the csp1 gene encoding PS1, one of the two major secreted proteins of Corynebacterium glutamicum: The deduced N-terminal region of PS1 is similar to the Mycobacterium antigen 85 complex," Mal. Microbiol., 6(16):2349-2362 (1992)
X66112	glt	Citrate syn	hase	Eikmanns, B.J. et al. "Cloning sequence, expression and transcriptional analysis of the Corynebacterium glutamicum gltA gene encoding citrate synthase," <i>Microbiol</i> , 140:1817-1828 (1994)
X67737	dapB	Dihydrodip	colinate reductase	
X69103	csp2	Surface lay	er protein PS2	Peyret, J.L. et al. "Characterization of the cspB gene encoding PS2, an ordered surface-layer protein in Corynebacterium glutamicum," <i>Mol. Microbiol.</i> , 9(1):97-109 (1993)
X69104		IS3 related	nsertion element	Bonamy, C. et al. "Identification of IS1206, a Corynebacterium glutamicum IS3-related insertion sequence and phylogenetic analysis," <i>Mol. Microbiol.</i> , 14(3):571-581 (1994)

Flack, M. et al. Learner perfect of leaf, and effect of leaf institution on bysics of concentration of the con			Table 2 (continued)	ned)
icd Isocitrate dehydrogenase (NADP+) GDHA Glutamate dehydrogenase (NADP+) recA aceA; thiX ATPase beta-subunit tuf Elongation factor Tu recA ATPase beta-subunit Inf Blongation factor Tu Inf Blongatio	X70959	leuA		Patek, M. et al. "Leucine synthesis in Corynebacterium glutamicum: enzyme activities, structure of leuA, and effect of leuA inactivation on lysine synthesis," <i>Appl. Environ. Microbiol.</i> , 60(1):133-140 (1994)
mtrA recA aceA; thiX recA tuf Elongation factor Tu recA aceB Malate synthase aceB Malate synthase 16S rDNA I Glutamate uptake system gluA; gluB; gluC; gluD dapE Succinyldiaminopimelate desuccinylase	X71489	icd	Isocitrate dehydrogenase (NADP+)	Eikmanns, B.J. et al. "Cloning sequence analysis, expression, and inactivation of the Corynebacterium glutamicum icd gene encoding isocitrate dehydrogenase and biochemical characterization of the enzyme," J. Bacteriol. 177(3):774-782 (1995)
recA aceA; thiX recA Tuf Elongation factor Tu recA aceB Malate synthase 16S rDNA I Glutamate uptake system gluA; gluB; gluC; gluD dapE Succinyldiaminopimelate desuccinylase	X72855	GDHA	Glutamate dehydrogenase (NADP+)	
aceA, thiX Partial Isocitrate lyase; ? ATPase beta-subunit tuf Elongation factor Tu recA aceB Malate synthase 16S ribosomal RNA I6S ribosomal RNA I6S ribosomal RNA Idea system gluD dapE Succinyldiaminopimelate desuccinylase	X75083, X70584	mtrA	5-methyltryptophan resistance	Heery, D.M. et al. "A sequence from a tryptophan-hyperproducing strain of Corynebacterium glutamicum encoding resistance to 5-methyltryptophan," <i>Biochem. Biophys. Res. Commun.</i> , 201(3):1255-1262 (1994)
aceA; thiX ATPase beta-subunit tuf Elongation factor Tu aceB Malate synthase 16S rDNA 16S ribosomal RNA 16S lutamate uptake system gluA; gluB; gluC; gluD Succinyldiaminopimelate desuccinylase	X75085	recA		Fitzpatrick, R. et al. "Construction and characterization of recA mulant strains of Corynebacterium glutamicum and Brevibacterium lactofermentum," <i>Appl. Microbiol Biotechnol.</i> , 42(4):575-580 (1994)
tuf recA aceB Malate synthase 16S ribosomal RNA gluA; gluB; gluC; gluD gluD Succinyldiaminopimelate desuccinylase	X75504	aceA; thiX	Partial Isocitrate lyase; ?	Reinscheid, D.J. et al. "Characterization of the isocitrate lyase gene from Corynebacterium glutamicum and biochemical analysis of the enzyme," J. Bacteriol., 176(12):3474-3483 (1994)
rec.A aceB Malate synthase 16S rDNA IGS ribosomal RNA gluA; gluB; gluC; gluD gluD Automate uptake system gluD Succinyldiaminopimelate desuccinylase	X76875		A TPase beta-subunit	Ludwig, W. et al. "Phylogenetic relationships of bacteria based on comparative sequence analysis of elongation factor Tu and ATP-synthase beta-subunit genes," <i>Antonie Van Lecuwenhoek</i> , 64:285-305 (1993)
aceB Malate synthase 16S rDNA 16S ribosomal RNA gluA; gluB; gluC; Glutamate uptake system gluD dapE Succinyldiaminopimelate desuccinylase	X77034	tuf		Ludwig, W. et al. "Phylogenetic relationships of bacteria based on comparative sequence analysis of elongation factor Tu and ATP-synthase beta-subunit genes," Antonie Van Leeuwenhoek, 64:285-305 (1993)
aceB Malate synthase 16S rDNA 16S ribosomal RNA gluA; gluB; gluC; Glutamate uptake system gluD dapE Succinyldiaminopimelate desuccinylase	X77384	recA		Billman-Jacobe, H. "Nucleotide sequence of a recA gene from Corynebacterium glutamicum," DNA Seq., 4(6):403-404 (1994)
16S ribosomal RNA gluA; gluB; gluC; Glutamate uptake system gluD dapE Succinyldiaminopimelate desuccinylase	X78491	aceB	Malate synthase	Reinscheid, D.J. et al. "Malate synthase from Corynebacterium giudinicum pta-ack operon encoding phosphotransacetylase: sequence analysis," <i>Microbiology</i> , 140:3099-3108 (1994)
gluA; gluB; gluC; Glutamate uptake system gluD dapE Succinyldiaminopimelate desuccinylase	X80629	16S rDNA	16S ribosomal RNA	Rainey, F.A. et al. "Phylogenetic analysis of the genera Knodococcus and Norcardia and evidence for the evolutionary origin of the genus Norcardia from within the radiation of Rhodococcus species," <i>Microbiol.</i> , 141:523-528 (1995)
dapE Succinyldiaminopimelate desuccinylase	X81191	gluA; gluB; gluC; gluD		Kronemeyer, W. et al. "Structure of the gluABCD cluster encoding the glutamate uptake system of Corynebacterium glutamicum," J. Bacteriol., 177(5):1152-1158 (1995)
	X81379	дарЕ	Succinyldiaminopimelate desuccinylase	Wehrmann, A. et al. "Analysis of different DINA fragments of Corynebacterium glutamicum complementing dapE of Escherichia coli," Microbiology, 40:3349-56 (1994)

			Table 2 (continued)	(panel)
X82061	16S rDNA	16S riboso	nal RNA	Ruimy, R. et al. "Phylogeny of the genus Corynebacterium deduced from analyses of small-subunit ribosomal DNA sequences," Int. J. Syst. Bacteriol., 45(4).740-746 (1995)
X82928	asd; lysC	Aspartate-	emialdchyde dehydrogenase; ?	Serebrijski, I. et al. "Multicopy suppression by asd gene and osmotic stress-dependent complementation by heterologous proA in proA mutants," J. Bacteriol., 177(24):7255-7260 (1995)
X82929	proA	Gamma-g	utamyl phosphate reductase	Serebrijski, I. et al. "Multicopy suppression by asd gene and osmotic stress-dependent complementation by heterologous proA in proA mutants," J. Bacteriol., 177(24):7255-7260 (1995)
X84257	16S rDNA	16S riboso	nal RNA	Pascual, C. et al. "Phylogenetic analysis of the genus Corynebacterium based on 16S rRNA gene sequences," Int. J. Syst. Bacteriol., 45(4):724-728 (1995)
X85965	aroP; dapE	Aromatic a	mino acid permease; ?	Wehrmann et al. "Functional analysis of sequences adjacent to dapE of C. glutamicum proline reveals the presence of aroP, which encodes the aromatic amino acid transporter," J. Bacterial, 177(20):5991-5993 (1995)
X86157	argB; argC; argD; argF; argJ		mate kinase; N-acetyl-gamma- nosphate reductase; ine aminotransferase; omithine ransferase; glutamate N- erase	Sakanyan, V. et al. "Genes and enzymes of the acetyl cycle of arginine biosynthesis in Corynebacterium glutamicum: enzyme evolution in the early steps of the arginine pathway," <i>Microbiology</i> , 142:99-108 (1996)
X89084	pia; ackA	Phosphate	cetyltransferase; acetate kinase	Reinscheid, D.J. et al. "Cloning, sequence analysis, expression and inactivation of the Corynebacterium glutamicum pta-ack operon encoding phosphotransacetylase and acetate kinase," <i>Microbiology</i> , 145:503-513 (1999)
X89850	attB	Attachmen	site	Le Marrec, C. et al. "Genetic characterization of site-specific integration functions of phi AAU2 infecting "Arthrobacter aureus C70," J. Bacteriol., 178(7):1996-2004 (1996)
X90356		Promoter fi	agment F1	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)
X90357		Promoter fi		Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> ; 142:1297-1309 (1996)
X90358		Promoter fr	agment F10	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> ; 142:1297-1309 (1996)
X90359		Promoter fr	ngment F13	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)

		Table 2 (continued	(pani
X90360		Promoter fragment F22	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)
X90361		Promoter fragment F34	Patek, M. et al. "Promoters from Coryncbacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)
X90362		Promoter fragment F37	Patek, M. et al. "Promoters from C. glutamicum: cloning, molecular analysis and search for a consensus motif," Microbiology, 142:1297-1309 (1996)
X90363		Promoter fragment F45	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)
X90364		Promoter fragment F64	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cloning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)
X90365		Promoter fragment F75	Patek, M. et al. "Promoters from Corynebacterium glutamicum. Cioniug, molecular analysis and search for a consensus motif," Microbiology, 142:1297-1309 (1996)
X90366		Promoter fragment PF101	Patek, M. et al. "Promoters from Corynebacterium glutameum. croning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)
X90367		Promoter fragment PF104	Patek, M. et al. "Promoters from Corynebacterium glutamicum. croning, molecular analysis and search for a consensus motif," Microbiology, 142:1297-1309 (1996)
X90368		Promoter fragment PF109	Patek, M. et al. "Promoters from Corynebacterium glutamicum: cioning, molecular analysis and search for a consensus motif," <i>Microbiology</i> , 142:1297-1309 (1996)
X93513	amt	Ammonium transport system	Siewe, R.M. et al. "Functional and genetic characterization of the (men.)" ammonium uptake carrier of Corynebacterium glutamicum," J. Biol. Chem., 271(10):5398-5403 (1996)
X93514	betP	Glycine betaine transport system	Corynebacterium glutamicum betP gene, encoding the transport system for the compatible solute glycine betaine," <i>J. Bacteriol.</i> , 178(17):5229-5234 (1996)
X95649	orf4		dapA-ORF4 operon of Corynebacterium glutamicum, encoding two enzymes involved in L-lysine synthesis," <i>Biotechnol. Lett.</i> , 19:1113-1117 (1997)
X96471	lysE; lysG	Lysine exporter protein; Lysine export regulator protein	Vrljuc, M. et al. "A new type of transporter with a new type of centural function: L-lysine export from Corynebacterium glutamicum," Mol. Microbiol., 22(5):815-826 (1996)

			Table 2 (continued)	(pənu
X96580	panB; panC; xylB	3-methyl-2. hydroxyme alanine liga	3-methyl-2-oxobutanoate hydroxyme hyltransferase; pantoate-beta- alanine ligace; xylulokinase	Sahm, H. et al. "D-pantothenate synthesis in Corynebacterium glutamicum and use of panBC and genes encoding L-valine synthesis for D-pantothenate overproduction," <i>Appl. Environ. Microbiol.</i> , 65(5):1973-1979 (1999)
X96962		Insertion se	sequence 1S1207 and transposase	
X99289		Elongation	actor P	Ramos, A. et al. "Cloning, sequencing and expression of the gene encoding elongation factor P in the amino-acid producer Brevibacterium lactofermentum (Corynebacterium glutamicum ATCC 13869)," Gene, 198 217-222 (1997)
Y00140	thrB	Homoserine kinase	kinase	Mateos, L.M. et al. "Nucleotide sequence of the homoserine kinase (thrB) gene of the Brevibacterium lactofermentum," <i>Nucleic Acids Res.</i> , 15(9):3922 (1987)
Y00151	qpp	Meso-diam (EC 1.4.1.1	n nopimelate D-dehydrogenase .1)	Ishino, S. et al. "Nucleotide sequence of the meso-diaminopimelate I)-dehydrogenase gene from Corynebacterium glutamicum," <i>Nucleic Acids Res.</i> , 15(9):3917 (1987)
Y00476	thrA	Homoserine	Homoserine dehydrogenase	Mateos, L.M. et al. "Nucleotide sequence of the homoserine dehydrogenase (thrA) gene of the Brevibacterium lactofermentum," <i>Nucleic Acids Res.</i> , 15(24):10598 (1987)
Y00546	hom; thrB	Homoserine kinase	ne dehydrogenase; homoserine	Peoples, O.P. et al. "Nucleotide sequence and fine structural analysis of the Corynebacterium glutamicum hom-thrB operon," <i>Mol. Microbiol.</i> , 2(1):63-72 (1988)
Y08964	murC, ftsQ/divD; ftsZ	UPD-N-ace division init protein; cell	UPD-N-ace ylnuramate-alanine ligase; division initiation protein or cell division protein; cell division protein	Honrubia, M.P. et al. "Identification, characterization, and chromosomal organization of the ftsZ gene from Brevibacterium lactofermentum," Mol. Gen. Genet., 259(1):97-104 (1998)
Y09163	putP	High affinit	High affinity proline transport system	Peter, H. et al. "Isolation of the putP gene of Corynebacterium glutamicumproline and characterization of a low-affinity uptake system for compatible solutes," <i>Arch. Microbiol.</i> , 168(2):143-151 (1997)
Y09548	pyc	Pyruvate ca	ca boxylase	Peters-Wendisch, P.G. et al. "Pyruvate carboxylase from Corynebacterium glutamicum: characterization, expression and inactivation of the pyc gene," <i>Microbiology</i> , 144:915-927 (1998)
Y09578	leuB	3-isopropyl	3-isopropylnalate dehydrogenase	Patek, M. et al. "Analysis of the leuB gene from Corynebacterium glutamicum," Appl. Microbiol. Biotechnol., 50(1):42-47 (1998)
Y12472		Attachment	site bacteriophage Phi-16	Moreau, S. et al. "Site-specific integration of corynephage Phi-16: The construction of an integration vector," <i>Microbiol.</i> , 145:539-548 (1999)
Y12537	proP	Proline/ectc	Proline/ecto ne uptake system protein	Peter, H. et al. "Corynebacterium glutamicum is equipped with four secondary carriers for compatible solutes: Identification, sequencing, and characterization of the proline/ectoine uptake system, ProP, and the ectoine/proline/glycine betaine carrier, EctP," J. Bacteriol, 180(22):6005-6012 (1998)

		Table 2 (continued)	nued)
Y13221	glnA	Glutamine synthetase I	Jakoby, M. et al. "Isolation of Corynebacterium glutamicum glnA gene encoding glutamine synthetase I," FEMS Microbiol. Lett., 154(1):81-88 (1997)
Y16642	pdl	Dihydrolipoamide dehydrogenase	
Y18059		Attachment site Corynephage 304L	Moreau, S. et al. "Analysis of the integration functions of φ,3041.: An integrase module among corynephages," <i>Virology</i> , 255(1):150-159 (1999)
721501	argS; lysA	Arginyl-tRNA synthetase; diaminopimelate decarboxylase (partial)	Oguiza, J.A. et al. "A gene encoding arginyl-tRNA synthetase is located in the upstream region of the lysA gene in Brevibacterium lactofermentum: Regulation of argS-lysA cluster expression by arginine," J. Bacteriol, 175(22):7356-7362 (1993)
221502	dapA; dapB	Dihydrodipicolinate synthase; dihydrodipicolinate reductase	Pisabarro, A. et al. "A cluster of three genes (dapA, orf2, and dapB) of Brevibacterium lactofermentum encodes dihydrodipicolinate reductase, and a third polypeptide of unknown function," J. Bacteriol., 175(9):2743-2749 (1993)
229563	thrC	Threonine synthase	Malumbres, M. et al. "Analysis and expression of the thrC gene of the encoded threonine synthase," <i>Appl. Environ. Microbiol.</i> , 60(7)2209-2219 (1994)
Z46753	16S rDNA	Gene for 16S ribosomal RNA	
249822	sigA	SigA sigma factor	Oguiza, J.A. et al "Multiple sigma factor genes in Brevibacterium lactofermentum: Characterization of sigA and sigB," J. Bacteriol., 178(2):550-553 (1996)
249823	galE; dtxR	Catalytic activity UDP-galactose 4- epimerase; diphtheria toxin regulatory protein	Oguiza, J.A. et al "The galE gene encoding the UDP-galactose 4-epimerase of Brevibacterium lactofermentum is coupled transcriptionally to the dmdR gene," <i>Gene</i> , 177:103-107 (1996)
249824	orfl; sigB	?; SigB sigma factor	Oguiza, J.A. et al "Multiple sigma factor genes in Brevibacterium lactofermentum: Characterization of sigA and sigB," J. Bacteriol., 178(2):550-553 (1996)
266534		Transposase	Correia, A. et al. "Cloning and characterization of an IS-like element present in the genome of Brevibacterium lactofermentum ATCC 13869," <i>Gene</i> , 170(1):91-94 (1996)
A sequence for the published v	A sequence for this gene was published in the indicat the published version. It is believed that the published	n the indicated reference. However, the sequence published version relied on an incorrect start or	A sequence for this gene was published in the indicated reference. However, the sequence obtained by the inventors of the present application is significantly longer than he published version. It is believed that the published version relied on an incorrect start codon, and thus represents only a fragment of the actual coding region.

TABLE 3: Corynebacterium and Brevibacterium Strains Which May be Used in the Practice of the Invention

Genus	species	ATCC	FERM	NRRL	CECT	NCIMB	CBS	NCTC	DSMZ
Brevibacterium	ammoniagenes	21054	1						
Brevibacterium	ammoniagenes	19350						1	
Brevibacterium	ammoniagenes	19351			<u> </u>		-		
Brevibacterium	ammoniagenes	19352							
Brevibacterium	ammoniagenes	19353			 				
Brevibacterium	ammoniagenes	19354							
Brevibacterium	ammoniagenes	19355			1				
Brevibacterium	ammoniagenes	19356							
Brevibacterium	ammoniagenes	21055	1					1	
Brevibacterium	ammoniagenes	21077						1	
Brevibacterium	ammoniagenes	21553	 		1				
Brevibacterium	ammoniagenes	21580	 		1				
Brevibacterium	ammoniagenes	39101							
Brevibacterium	butanicum	21196							
Brevibacterium	divaricatum	21792	P928						
Brevibacterium	flavum	21474					-		
Brevibacterium	flavum	21129		ĺ					
Brevibacterium	flavum	21518						Ī	
Brevibacterium	flavum			B11474					
Brevibacterium	flavum			B11472				Í	
Brevibacterium	flavum	21127							
Brevibacterium	flavum	21128							
Brevibacterium	flavum	21427					_		
Brevibacterium	flavum	21475							
Brevibacterium	flavum	21517							
Brevibacterium	flavum	21528							
Brevibacterium	flavum	21529					i 		
Brevibacterium	flavum			B11477	_				ļ
Brevibacterium	flavum			B11478	<u> </u>				
Brevibacterium	flavum	21127			<u> </u>				
Brevibacterium	flavum			B11474				ļ	
Brevibacterium	healii	15527						<u> </u>	ļ
Brevibacterium	ketoglutamicum	21004			<u> </u>				
Brevibacterium	ketoglutamicum	21089			<u> </u>				
Brevibacterium	ketosoreductum	21914							ļ
Brevibacterium	lactofermentum				70			<u> </u>	ļ
Brevibacterium	lactofermentum				74				
Brevibacterium	lactofermentum				77				
Brevibacterium	lactofermentum	21798						1	
Brevibacterium	lactofermentum	21799						<u> </u>	L
Brevibacterium	lactofermentum	21800						1	L
Brevibacterium	lactofermentum	21801							
Brevibacterium	lactofermentum			B11470					
Brevibacterium	lactofermentum			B11471					

Genus	species	ATCC	FERM	NRRL	CECT	NCIMB	CBS	NCTC	DOMIZ
Brevibacterium	lactofermentum	21086					······································	ļ	
Brevibacterium	lactofermentum	21420				<u> </u>			
Brevibacterium	lactofermentum	21086							<u> </u>
Brevibacterium	lactofermentum	31269							
Brevibacterium	linens	9174							
Brevibacterium	linens	19391							
Brevibacterium	linens	8377							
Brevibacterium	paraffinolyticum					11160			ļ
Brevibacterium	spec.						717.73		<u> </u>
Brevibacterium	spec.						717.73	L	
Brevibacterium	spec.	14604							
Brevibacterium	spec.	21860							
Brevibacterium	spec.	21864						<u> </u>	
Brevibacterium	spec.	21865							
Brevibacterium	spec.	21866						<u> </u>	ļ
Brevibacterium	spec.	19240							ļ
Corynebacterium	acetoacidophilum	21476							ļ
Corynebacterium	acetoacidophilum	13870							
Corynebacterium	acetoglutamicum			B11473				ļ	
Corynebacterium	acetoglutamicum			B11475	5			ļ	
Corynebacterium	acetoglutamicum	15806							
Corynebacterium	acetoglutamicum	21491							
Corynebacterium	acetoglutamicum	31270							
Corynebacterium	acetophilum			B3671					
Corynebacterium	ammoniagenes	6872						2399	
Corynebacterium	ammoniagenes	15511							
Corynebacterium	fujiokense	21496							
Corynebacterium	glutamicum	14067							
Corynebacterium	glutamicum	39137							
Corynebacterium	glutamicum	21254							
Corynebacterium	glutamicum	21255					<u> </u>		↓
Corynebacterium	glutamicum	31830				J			
Corynebacterium	glutamicum	13032							
Corynebacterium	glutamicum	14305							
Corynebacterium	glutamicum	15455					<u> </u>		_
Corynebacterium	glutamicum	13058							
Corynebacterium	glutamicum	13059							
Corynebacterium	glutamicum	13060							
Corynebacterium	glutamicum	21492							
Corynebacterium	glutamicum	21513							<u> </u>
Corynebacterium	glutamicum	21526							
Corynebacterium	glutamicum	21543							
Corynebacterium	glutamicum	13287		1					
Corynebacterium	glutamicum	21851		1					
Corynebacterium	glutamicum	21253	+-		1				
Corynebacterium	glutamicum	21514			-				
Corynebacterium	glutamicum	21516							
Corynebacterium	glutamicum	21299			1				

Genus	species	ATCC	FERM	NRRL	CECT	NCIMB	CBS	NCTC	DSMZ
Corynebacterium	glutamicum	21300							
Corynebacterium	glutamicum	39684							
Corynebacterium	glutamicum	21488							
Corynebacterium	glutamicum	21649			†			1	
Corynebacterium	glutamicum	21650	1			†		1	
Corynebacterium	glutamicum	19223	 		1			<u> </u>	
Corynebacterium	glutamicum	13869	<u> </u>		1				
Corynebacterium	glutamicum	21157							
Corynebacterium	glutamicum	21158			1			1	
Corynebacterium	glutamicum	21159			†	<u> </u>			
Corynebacterium	glutamicum	21355				†	- 13	1	
Corynebacterium	glutamicum	31808							
Corynebacterium	glutamicum	21674			1		_		
Corynebacterium	glutamicum	21562		·	† <u>-</u>				
Corynebacterium	glutamicum	21563			1				
Corynebacterium	glutamicum	21564			<u> </u>			1	
Corynebacterium	glutamicum	21565	ļ -		1				
Corynebacterium	glutamicum	21566						1	
Corynebacterium	glutamicum	21567							
Corynebacterium	glutamicum	21568	†		1			1	
Corynebacterium	glutamicum	21569	†	-					
Corynebacterium	glutamicum	21570	1		<u> </u>				
Corynebacterium	glutamicum	21571							
Corynebacterium	glutamicum	21572							
Corynebacterium	glutamicum	21573							
Corynebacterium	glutamicum	21579							
Corynebacterium	glutamicum	19049							
Corynebacterium	glutamicum	19050			1				
Corynebacterium	glutamicum	19051							
Corynebacterium	glutamicum	19052							
Corynebacterium	glutamicum	19053							
Corynebacterium	glutamicum	19054			I				
Corynebacterium	glutamicum	19055							
Corynebacterium	glutamicum	19056							
Corynebacterium	glutamicum	19057							
Corynebacterium	glutamicum	19058							
Corynebacterium	glutamicum	19059							
Corynebacterium	glutamicum	19060							
Corynebacterium	glutamicum	19185							
Corynebacterium	glutamicum	13286						<u> </u>	
Corynebacterium	glutamicum	21515							
Corynebacterium	glutamicum	21527							
Corynebacterium	glutamicum	21544							
Corynebacterium	glutamicum	21492							
Corynebacterium	glutamicum			B8183					
Corynebacterium	glutamicum			B8182					
Corynebacterium	glutamicum			B12416					
Corynebacterium	glutamicum			B12417					

Genus	species	ATCC.	FERM	NRRL	CECT	NCIMB	CBS	NCTC	DSMZ
Corvnebacterium	glutamicum			B12418					
Corynebacterium	glutamicum			B11476					
Corvnebacterium	glutamicum	21608							
Corynebacterium	lilium		P973						
Corynebacterium	nitrilophilus	21419				11594			
Corynebacterium	spec.		P4445					ļ	ļ
Corynebacterium	spec.		P4446						ļ
Corynebacterium	spec.	31088			<u> </u>				ļ
Corynebacterium	spec.	31089						J	
Corynebacterium	spec.	31090							
Corynebacterium	spec.	31090				ļ			ļ
Corynebacterium	spec.	31090				ļ			20145
Corynebacterium	spec.	15954		<u> </u>					20145
Corynebacterium	spec.	21857							<u> </u>
Corynebacterium	spec.	21862			ļ				-
Corynebacterium	spec.	21863				1	L		<u> </u>

ATCC: American Type Culture Collection, Rockville, MD, USA

FERM: Fermentation Research Institute, Chiba, Japan

NRRL: ARS Culture Collection, Northern Regional Research Laboratory, Peoria, IL, USA

CECT: Coleccion Espanola de Cultivos Tipo, Valencia, Spain

NCIMB: National Collection of Industrial and Marine Bacteria Ltd., Aberdeen, UK

CBS: Centraalbureau voor Schimmelcultures, Baarn, NL

NCTC: National Collection of Type Cultures, London, UK

DSMZ: Deutsche Sammlung von Mikroorganismen und Zellkulturen, Braunschweig, Germany

For reference see Sugawara, H. et al. (1993) World directory of collections of cultures of microorganisms: Bacteria, fungi and yeasts (4th edn), World federation for culture collections world data center on microorganisms, Saimata, Japen.

					_				95											
	Date of Deposit	29-Jun-99	29-Jun-99	08-OCT-	1997 (Rel. 52, Created)	07-OCT- 1996	17-DEC- 1993	28-Jul-99	2-Aug-99	2-Aug-99		17-Jun-98	14-Jan-97	12-Nov-98	20-Aug-97	9-Sep-99	29-Sep-99	30-MAR- 1999	2-Sep-99	2-Sep-99
	% homology (GAP)	40,956	40,956	42,979		42,979	39,097	95,429	31,111	31,111		37,753	35,669	35,669	42,896	40,210	41,176	36,783	40,296	40,296
	Source of Genbank Hit	Lycopersicon esculentum	Lycopersicon esculentum	Corynebacterium	giutamicum	Unknown.	Escherichia coli	Corynebacterium glutamicum	Drosophila melanogaster	Drosophila melanogaster		Mycobacterium tuberculosis	Escherichia coli	Escherichia coli	Homo sapiens	Corynebacterium diphtheriae	Unknown.	Homo sapiens	Homo sapiens	Homo sapiens
Table 4: Alignment Results	Name of Genbank Hit	EST257217 tomato resistant, Cornell Lycopersicon esculentum cDNA clone cLER17D3, mRNA sequence.	EST257217 tomato resistant, Cornell Lycopersicon esculentum cDNA clone cLER17D3, mRNA sequence.	Base sequence of sucrase gene.		Sequence 4 from patent US 5556776.	E. coli chromosomal region from 89.2 to 92.8 minutes.	gDNA encoding aspartate transferase (AAT).	Drosophila melanogaster chromosome 3 clone BACR02003 (D797) RPCI-98 02:0.3 map 998-998 strain y; cn bw sp, *** SEQUENCING IN PROGRESS ***, 113 unordered pieces.			Mycobacterium tuberculosis H37Rv complete genome; segment 122/162.	Escherichia coli K-12 genome, approximately 63 to 64 minutes.		ng83f04.s1 NCI_CGAP_Pr6 Homo sapiens cDNA clone IMAGE:941407 similar to SW.DYR_LACCA P00381 DIHYDROFOLATE REDUCTASE ;; mRNA sequence.) Homo sapiens chromosome 17, clone hRPK 472_J_18, complete sequence.		IN PROGRESS ***, 93 unordered pieces. Nomo sapiens chromosome 19 clone CIT-HSPC_490E21, *** SEQUENCING Homo sapiens IN PROGRESS ***, 93 unordered pieces.
	Length Access on	AI776129	AI776129	E11760		126124	00000	E16763	AC007892	AC007892		AL008957	U29581	AE000356	AA494237	AF161327	AR041189	AC007110	AC008537	AC008337
	Length	483	483	6911		6911	176195 U00006	2517	134257	134257		56414	71128	10405	367	2021	654	148336	170030	170030
	Genbank Hit	GB_EST33:AI776129	GB_EST33.AI776129	EM_PAT:E11760		GB_PAT:126124	GB_BA2:ECOUW89	GB_PAT:E16763	GB_HTG2.AC007892 134257 AC007892	GB_HTG2.AC007892 134257 AC007892		GB_BA1:MTV002	GB_BA1.ECU29581	GB_BA2:AE000366 1040	GB_EST15:AA494237	GB_BA2:AF161327	GB_PAT:AR041189	GB_PR4:AC007110	GB_HTG3.AC008537 170030 AC008537	GB_HTG3.AC008537 170030 AC008537
	length (NT)	3579		1059				1401				798			579			1170		
	# Q	rxa00023		rxa00044				rxa00064			rxa00072	rxa00105			rxa00106			xa00115		

	19-0C1-	07-OCT. 1996	8-Apr-99	17-Jun-98	15-Jun-99 17-Jun-98	17-Jun-98	31-OCT- 1996	22-Nov-99	18-Jun-98	26-Jul-93	29-Apr-97	18-Jun-98	17-Jun-98	03-DEC- 1996	18-Jun-98	15-Jun-96	23-DEC- 1996	10-Feb-99
,	36,235	36,821	38,124	43,571	41,116 39,726	36,788	61,914	51,325	63,365	56,080	47,514	60,714	39,229	36,618	61,527	59,538	55,396	52,666
	Caulobacter crescentus	Unknown.	Oryza sativa	Mycobacterium tuberculosis	Streptomyces argillaceus Mycobacterium tuberculosis	Mycobacterium tuberculosis	Trichomonas vaginalis	Drosophila melanogaster	Mycobacterium tuberculosis	Pseudomonas aeruginosa	Lactobacillus leichmannii	Mycobacterium tuberculosis	Mycobacterium tuberculosis	Mycobacterium tuberculosis	Mycobacterium tuberculosis	Mycobacterium leprae	Pseudomonas aeruginosa	Streptomyces coelicolor
Table 4 (continued)	Caulobacter crescentus Sst1 (sst1), S-layer protein subunit (rsaA), ABC transporter (rsaD), membrane forming unit (rsaE), putative GDP-mannose-4,6 dehydratase (lpsA), putative acetyltransferase (lpsB), putative perosamine synthetase (lpsC), putative mannosyltransferase (lpsD), putative mannosyltransferase (lpsE), and putative perosamine transferase (lpsE) genes, complete cds.	Sequence 6 from patent US 5500353.	nbxb0062D16r CUGI Rice BAC Library Oryza sativa genomic clone nbxb0062D16r, genomic survey sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 139/162.	Streptomyces argillaceus mithramycin biosynthetic genes. Mycobacterium tuberculosis H37Rv complete genome, segment 139/162.	Mycobacterium tuberculosis H37Rv complete genome, segment 139/162.	Trichomonas vaginalis S-adenosyl-L-homocysteine hydrolase gene, complete cds.	Drosophila melanogaster chromosome X clone BACR36D15 (D887) RPCI-98 36.D.15 map 13C-13E strain y; cn bw sp, *** SEQUENCING IN PROGRESS ***, 74 unordered pieces.	Mycobacterium tuberculosis H37Rv complete genome, segment 61/162.	Pseudomonas aeruginosa aspartate transcarbamoylase (pyrB) and dihydroorotase-like (pyrX) genes, complete cds's.	L.Ieichmannii pyrB gene.	Mycobacterium tuberculosis H37Rv complete genome; segment 61/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 121/162.	Mycobacterium tuberculosis sequence from clone y154.	Mycobacterium tuberculosis H37Rv complete genome; segment 61/162.	Mycobacterium leprae cosmid B937 DNA sequence.	Pseudomonas aeruginosa dihydrodipicolinate reductase (dapB) gene, partial cds, carbamoylphosphate synthetase small subunit (carA) and carbamoylphosphate synthetase large subunit (carB) genes, complete cds, and FtsJ homolog (ftsJ) gene, partial cds.	Streptomyces coelicolor cosmid 9B10.
	A F062345	118647	AQ446197	295121	AJ007932 Z95121	Z 95121	U40872	AC010706	281011	L19649	X84262	281011	298209	AD000002	281011	L78820	U81259	AL009204
	16458	3300	751	36330	15176 36330	36330	1882	169265	20431	2273	۱468	20431	13935	40221	20431	38914	7285	33320
	GB_BA2.AF062345	GB_PAT:118647	GB_GSS13;AQ44619_751 7	GB_BA1:MTY20B11	GB_BA1:SAR7932 GB_BA1:MTY20B11	GB_BA1:MTY20B11	GB_IN2:TVU40872	GB_HTG6.AC010706 169265 AC010706	GB_BA1.MTCY2B12	GB_BA1.PSEPYRBX	GB_BA1:LLPYRBDNA 1468	GB_BA1:MTCY2B12	GB_BA1.MTCY154	GB_BA1:MSGY154	GB_BA1:MTCY2B12	GB_BA1:MSGB937C S	GB_BA1:PAU81259	GB_BA1:SC9B10
	1284			732		1557			1059			1464			1302			1233
	rxa00116			rxa00131		rxa00132			rxa00145			rxa00146			rxa00147			rxa00156

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	26-MAR- 1998	6-Feb-99	21-Aug-99	21-Aug-99	16-0CT-	13-MAR-	1999	13-MAK-	1999 31-Jan-99	;	03-DEC- 1999	03-DEC-	1999	11-Nov-99		22-Nov-98	22-Nov-98	22-Nov-98	2-Aug-96	22-MAR-	1999 16 OCT	1999	18-MAY-	1999	18-Apr-98	26-Feb-99	30-Jan-92		
	54,191	46,667	37,451	37,451	38,627	92,113	0	93,702	34,221		37,965	37,965		38,796		38,227	38,227	38.227		40,135	703.00	720,86	98,237		36,616	37,095	100 000		
	Mycobacterium avium	Propionibacterium freudenreichii	Homo sapiens	Homo sapiens	Drosophila melanogaster	Corynebacterium	glutamicum	Corynepacterium	glutamicum Rattus sp		Homo sapiens	Homo sapiens		Homo sapiens		Homo sapiens	Homo sapiens	Homo sapiens	Enterobacter agglomerans	Rhodobacter capsulatus		Orosopiiila iiielanogastei	Conynebacterium	glutamicum		, Caenorhabditis elegans	Corynehacterium	glutamicum	
lable 4 (continued)	3 Mycobacterium avium strain GIR10 transcriptional regulator (mav81) gene, partial cds, aconitase (acn), invasin 1 (inv1), invasin 2 (inv2), transcriptional regulator (moxR), ketoacyl-reductase (fabG), enoyl-reductase (inhA) and ferrochelatase (mav272) genes, complete cds.	Propionibacterium freudenreichii hemY, hemH, hemB, hemX, hemR and hem! cenes, complete cds.			unordered pieces. 8 Drosophila melanogaster chromosome 3L/62B1 clone RPC198-10D15, *** SEDI IENCING IN PROGRESS *** 51 unordered nieces.						Homo sapiens chromosome 20 clone RP5-850E9, *** SEQUENCING IN PROGRESS ***, in unordered pieces.	Homo sapiens ch		 Human chromosome 14 DNA sequence *** IN PROGRESS *** BAC R-412H8 Homo sapiens of RPCI-11 library from chromosome 14 of Homo sapiens (Human) complete 	Sequence	9 Homo sapiens clone RG252P22, *** SEQUENCING IN PROGRESS ***, 3	unordered pieces. 9 Homo sapiens clone RG252P22, *** SEQUENCING IN PROGRESS ***, 3		Plasmid pEA3 nitrogen fixation genes.	4 Rhodobacter capsulatus molybdenum cofactor biosynthetic gene cluster,	partial sequence.	SEQUENCING IN PROGRESS *** 64 unordered pieces.	8 Corynebacterium glutamicum 3-dehydroquinase (aroD) and shikimate	dehydrogenase (aroE) genes, complete cds.		7 Caenorhabditis elegans clone Y76B12, *** SEQUENCING IN PROGRESS ***	25 unordered pieces. C olutamicum Ivst gene for t-Ivsine permease		
	AF002 33	D85417	AC008 67	1C008 67	AC010118	AB024708	- 1	ABU24 /U8	AI2327 2		NL12176	AL1217		VL1217		AC00507	AC0050	AC0050	X99694	AF1284	,0,00		AF1245		AC0045	AC0069	X60312		
	15437 A	7984 C	174223 #	174223 AC008	80605	8734 A		0/34	528		117353 7	117353 4		159400 /		110000 /	110000 /	110000 /	_	2477	1,00000	006001	1758 #			188972 #	4232		
	GB_BA2.AF002133	GB_BA1:D85417	GB_HTG3.AC008167 174223 AC008	GB_HTG3.AC008167	GB_HTG4:AC010118	GB_BA1:AB024708	900000000000000000000000000000000000000	0B_BA1.AB024/08	GB_EST24:AI232702		GB_HTG2.HSDJ850E 117353 AL121768 9	GB_HTG2:HSDJ850E 117353 AL121758	6	GB_PR2:CNS01DSA 159400 AL121766		GB_HTG2.AC005079 110000 AC00507	_0 GB_HTG2:AC005079_110000_AC005079	_1 _B HTG2.AC005079 110000 AC0050	 GB_BA1:PPEA3NIF	GB_BA2:AF128444	44.0000 A. A.O.F.D. G.O.	1101000 X 2011 - a0	GB_BA2.AF124518		GB_PR3.AC004593	GB_HTG2:AC006907	GB BA1-CGLYSI		

rxa00216 1113

rxa00219 1065

rxa00223 1212

xa00229 803

rxa00241 1626

rxa00166 783

rxa00198 672

Table 4 (continued)

WC	01/0	0843									98	3								F	CT	71 B	00,	(009	23			
11-Aug-99	11-Aug-99	23-MAY-	1997 23-MAY- 1997	9-Feb-99	08-OC1- 1997 (Rel	52, Created) 29-Sep-97	6-Jan-98	9-Apr-97	20-840-96	21-Nov-96		3-Feb-99	29-Sep-97		24-Jun-99	15-Jun-96	24-Jun-99		15-Jun-96	24-Jun-99	27-Anr-93	17-Jun-97		02-DEC- 1994	20-Sep-95		28-Aug-97	
34,947	34,947	36,496	37,544	41,856	34,741	34 741	36,943	36,658	38 190	99,111		98,489	98,207		35,615	60,917	44,606		52,516	38,079	39 351	808,66		99,617	99,170		100,000	
Plasmodium falciparum	Plasmodium falciparum	Entamoeba histolytica	Entamoeba histolytica	Mus musculus	Bacillus sp.	Bacillis on	Caenorhabditis elegans	Corynebacterium	glutamicum	Corynebacterium	glutamicum	Corynebacterium	Corynebacterium	glutamicum	Mycobacterium tuberculosis	Mycobacterium leprae	Mycobacterium	tuberculosis	Mycobacterium leprae	Mycobacterium	tuberculosis Bos taurus		glutamicum	Unknown.	Corynebacterium	glutamicum	Corynebacterium olutamicum	GIOCOLINICALINI
Table 4 (continued) Plasmodium falcipatum chromosome 13 strain 3D7, *** SEQUENCING IN	PROGRESS ***, in unordered pieces. Plasmodium falciparum chromosome 13 strain 3D7, *** SEQUENCING IN	PROGRESS ***, in unordered pieces. Entamoeba histolytica unconventional myosin IB mRNA, complete cds.	Entamoeba histolytica unconventional myosin IB mRNA, complete cds.	Mus musculus connexin-36 (Cx36) gene, complete cds.	DNA encoding precursor protein of alkaline cellulase.		gDNA encoding alkaline cellulase. Channihabditis alaqans cosmid KOSE6	Corynebacterium glutamicum multidrug resistance protein (cmr) gene,	complete cds.	Rattus norvegicus clone NZ/ mKNA. Comparactarium alutamicum biotin synthase (bioB) gene, complete cds.		Brevibacterium flavum gene for biotin synthetase, complete cds.	DNA seguence encoding Brevibacterium flavum biotin-synthase.		Mycobacterium tuberculosis H37Rv complete genome; segment 99/162.	Mycobacterium leprae cosmid B32 DNA sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 99/162.		Mycobacterium leprae cosmid B32 DNA sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 99/162		Bovine elastin a mkNA, complete cds. Corynebacterium glutamicum thrC gene for threonine synthase (EC 4.2.99.2).		Sequence 4 from Patent WO 8809819.	Bravibarterium (actofermentum: ATCC 13869:: DNA (genomic):		Corynebacterium glutamicum glnA gene.	
AI 049180		U89655	U89655	AF016190	E09719		E02133	U43535		U30789	03150	D14084	E03937		Z70692	L78818	270692		L78818	Z70692		J02717 X56037		109078	720563	2000	Y13221	
192581	192581	3219	3219	2939	3505		3494	2531		3510	<u> </u>	1647	1005	2	38110	36404	38110		36404	38110		3242))	3146	1802	700	3686	
CD LTC1.DEMAI 13D 192581	GB HTG1:PFMAI 13P 192581	1 GB IN2:EHU89655	GB_IN2.EHU89655	GB RO:AF016190	EM_PAT.E09719		GB_PAT:E02133	GB_BA1.CGU43535		GB_RO:RNU30789	GB_BAZ:CGU31201	GB_BA1:BRLBIOBA	OB DAT: E03037	10800 TAL 00	GB_BA1:MTCY427	GB_BA1:MSGB32CS	GB BA1:MTCY427		GB_BA1:MSGB32CS	GB BA1:MTCY427	•	GB_OM.BOVELA		GB_PAT:109078	Vondut Id-bad do	N	GB_BA1.CGGLNA	
		1197		531				351			1125				1461				3258			756	200				1554	
		rxa00262		990000	0000			rxa00278			rxa00295				rxa00323				rxa00324			000000	raduoso				rxa00335	

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		GB_BA2:AF005635	1690	AF005635	Table 4 (continued) Corynebacterium glutamicum glutamine synthetase (glnA) gene, complete	Corynebacterium	906'86	14-Jun-99
		GB BA1.MSGB27CS 38793	38793	L78817	cds. Mycobacterium leprae cosmid B27 DNA sequence.	glutamicum Mycobacterium leprae	66,345	15-Jun-96
rxa00347	108	GR EST27.AI455217 6	624	A1455217	1 D21828 3prime 1 D Drosophija mejanogaster embryo pOT2 Drosophija	Drosophila metanogaster	34 510	09-MAR-
					melanogaster cDNA clone LD21828 3prime, mRNA sequence.) 	1999
		GB_BA2:SSU30252 2	2891	U30252	Synechococcus PCC7942 nucleoside diphosphate kinase and ORF2 protein genes, complete cds, ORF1 protein gene, partial cds, and neutral site I for vector use.	Synechococcus PCC7942	37,084	29-OCT- 1999
		GB_EST21.AA911262 581	581	AA911252		Homo sapiens	37,500	21-Apr-98
rxa00351	1578	GB_BA1:MLU15187	36138	U15187	Mycobacterium leprae cosmid L296.	Mycobacterium leprae	52,972	09-MAR- 1995
		GB_IN2.AC004373	72722	AC004373	 Drosophila melanogaster DNA sequence (P1 DS05273 (D80)), complete sequence. 	Orosophila melanogaster	46,341	17-Jul-98
			3197	AF145653	3 Drosophila melanogaster clone GH08860 BcDNA.GH08860 (BcDNA.GH08860) mRNA, complete cds.	Drosophila melanogaster	49,471	14-Jun-99
rxa00365	727	GB_BA1:AB024708 8	8734	AB024708	 Corynebacterium glutamicum gltB and gltD genes for glutamine 2- oxoglutarate aminotransferase large and small subunits, complete cds. 	Corynebacterium glutamicum	96,556	13-MAR- 1999
		GB_BA1:MTCY1A6	37751	Z83864	Mycobacterium tuberculosis H37Rv complete genome; segment 159/162.	Mycobacterium tuberculosis	39,496	17-Jun-98
		GB_BA1:SC3A3	15901	AL1098-9	 Streptomyces coelicolor cosmid 3A3. 	Streptomyces coelicolor A3(2)	37,946	16-Aug-99
rxa00366	480	GB_BA1.AB024708	8734	AB0247 8	Corynebacterium glutamicum gltB and gltD genes for glutamine 2- coronitarate aminotransferase large and small subunits, complete rds	Corynebacterium	99,374	13-MAR- 1999
		GB_BA1:MTCY1A6	37751	Z83864	Mycobacterium tuberculosis H37Rv complete genome; segment 159/162.	gratarricarri Mycobacterium tuberculosis	41,333	17-Jun-98
		GB_BA1:SC3A3	15901	AL1098-9	Streptomyces coelicolor cosmid 3A3.	Streptomyces coelicolor A3(2)	37,554	16-Aug-99
rxa00367	4653	GB_BA1:AB024708	8734	AB0247		Corynebacterium	99,312	13-MAR-
		GB_BA1:MTCY1A6	37751	Z83864	oxoglutarate aminotransferase large and small subunits, complete cds. Mycobacterium tuberculosis H37Rv complete genome; segment 159/162.	glutamicum Mycobacterium	36,971	1999 17-Jun-98
		GB_BA1:SC3A3	15901	AL109849	Streptomyces coelicolor cosmid 3A3.	tuberculosis Streptomyces coelicolor A3(2)	37,905	16-Aug-99
rxa00371	1917	GB_VI:SBVORFS	7568	M89923	Sugarcane bacilliform virus ORF 1,2, and 3 DNA, complete cds.	Sugarcane baciliform virus 35,843	35,843	12-Jun-93
		GB_EST37.AI967505	380	A196750	Ljirnpest03-215-c10 Ljirnp Lambda HybriZap two-hybrid library Lotus japonicus cDNA clone LP215-03-c10 5' similar to 60S ribosomal protein L39, mRNA sequence.	Lotus japonicus	42,593	24-Aug-99
		GB_IN1.CELK09H9	37881	AF043700		Caenorhabditis elegans	34,295	22-Jan-98

WO 01	/008	343							10	0							PCT	[/ IB 0)/009	23
24-MAR- 1995 17-OCT-	1996	15-Jul-99	18-DEC- 1995	17-Jun-98	03-DEC- 1996	27-Aug-99 10-Jun-99	22-MAY- 1999	10-Sep-99	m	2-Aug-99		17-Jun-98	03-DEC- 1996	24-Jun-97	19-MAR- 1998	8-Jun-99	06-DEC-	19-MAR- 1998	23-Jun-99	31-Aug-98
36,832		36,728	54,175	61,143	61,143	43,981 35,44 4	34,821	40,472	38,586	38,509		36,308	39,282	39,228	99,672	40,830	50,161	99,920	52,898	37,565
Caulobacter crescentus	Emericella ricularis	Homo sapiens	Pseudomonas aeruginosa	Mycobacterium tuberculosis	Mycobacterium tuberculosis	Mycobacterium leprae Homo sapiens	Homo sapiens	Schistosoma mansoni	Unknown.	 Kaposi's sarcoma- associated herpesvirus 		Mycobacterium tuberculosis	Mycobacterium tuberculosis	Mycobacterium leprae	Corynebacterium	Corynebacterium	diphtheriae Pseudomonas alcaligenes	Corynebacterium	Mycobacterium tuberculosis	N-Onchocerca volvulus
Table 4 (continued) Caulobacter crescentus uroporphyrinogen decarboxylase homolog (hemE) gene, partial cds.	A.nidulans sD gene.	HS_5505_B1_C04_T7A RPCI-11 Human Male BAC Library Homo sapiens	genomic cone Prate Tool Col-7 Now 1, goronno con 2, col-7	Mycobacterium tuberculosis H37Rv complete genome; segment 28/162.	Mycobacterium tuberculosis sequence from clone y224.	Mycobacterium leprae cosmid B1222. Homo sapiens chromosome 17 clone hRPK.515_E_23 map 17, ***	SEQUENCING IN PROGRESS ***, 2 ordered pieces. Homo sapiens chromosome 17 clone hRPK.515_O_17 map 17, *** peroviewichio in papogaess *** 8 innordered pieces.		Schistosoma mansoni cDNA clone SMMAS14 5 end, mkiva sequence:	rus ORF 68 gene, partia clear antigen, ORF K14,	putative phosphoribosylformylglycinamidine synthase, and LAMP	(LAMP) genes, complete cus. Mycobacterium tuberculosis H37Rv complete genome; segment 28/162.	Mycobacterium tuberculosis sequence from clone y224.	Mycobarterium lenrae cosmid B1306 DNA.	Corynebacterium glutamicum homoserine O-acetyltransferase (metA) gene.	complete cds. Corynebacterium diphtheriae heme uptake locus, complete sequence.	Pseudomonas alcaligenes outer membrane Xcp-secretion system gene		complete cds. Mycobacterium tuberculosis H37Rv complete genome; segment 143/162.	SWOVAMCAQ02A05SK Onchocerca volvulus adult male cDNA (SAW98MLW-Onchocerca volvulus OvAM) Onchocerca volvulus cDNA clone SWOvAMCAQ02A05 5', mRNA sequence.
U13664	Y08866	AQ730303	X82072	Z95558	AD000004	AL049491 AC006269	AC007638	AW017053		AF148805		Z95558	AD000004	V13803	AF052652	AF109162	AF092918	AF052652	AL021841	Al111288
1678	1299	483	4444	40838	40051	34714 167171	178053	613	000	32207 28559		40838	40051	7763	2096	4514	20758	2096	53662	750
GB_BA1:CCU13664	GB_PL1:ANSDGENE	GB_GSS4:AQ730303	GB_BA1:PAHEML	GB_BA1:MTY25D10	GB_BA1:MSGY224	GB_BA1:MLCB1222 GB_HTG2:AC006269	GB_HTG2:AC007638 178053	GB_EST38:AW01705	3	GB_PAT:AR065852 GB_VI:AF148805		GB_BA1:MTY25D10	GB_BA1:MSGY224		GB_BA2:AF052652	GB BA2 AF109162	GB BA2.AF092918	GB_BA2:AF052652	GB_BA1:MTV016	GB_EST23.Al111288 750
1245			1425			1467		843				1017			623			1254		
rxa00377			rxa00382			rxa00383		rxa00391				rxa00393			rxa00402			rxa00403		

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												101																		
23-Jun-99	08-DEC-	1998 23-Jun-99	17-Jun-98	0	1996	15-Jun-96	27-Jul-94	12-0CT-	1999	12-OCT-	1999 18-Nov-98	5-Aug-98	18-Nov-98	23-Jun-99	00	17-Aug-99	ee-fin⊄-c	27-0CT-	1999	4-Jun-98	4-Jun-98	23-Nov-99	03.050	1999	03-DEC-	1999	26-Nov-98	16-0CT-	1999 16-OCT-	1999
57.259	34,179	40,169	62 031		206,10	39,651	38,677	36,335		36,335	31,738	43,262	37,647	37,088	0	45,538	0/7,04	43,080		42,931	36,702	38,027	34 521	10,10	34,521		56,410	34,959	24 050	6 6 7
Mycobacterium		Mycobacterium	tuberculosis Mycobacterium	tuberculosis	iviycobacterium tuberculosis	Mycobacterium leprae	Ralstonia eutropha	Homo sapiens		Homo sapiens	Homo sapiens	Streptomyces coelicolor	Homo sapiens	Mycobacterium	tuberculosis	Kumex acetosa		Streptomyces Iividans		Streptomyces coelicolor	Streptomyces coeficolor	Homo sapiens		200000000000000000000000000000000000000	Homo sapiens		Streptomyces coelicolor	Drosophila melanogaster	Drosophia melanosom	
Table 4 (continued) Mycobacterium tuberculosis H37Ry complete penome: segment 143/162	Homo sapiens Xp22-166-169 GSHB-523A23 (Genome Systems Human BAC	library) complete sequence. Mycobacterium fuberculosis H37Ry complete genome: segment 143/162.	Myrobacterium tuherrulosis H37Ry complete genome: segment 156/152		Mycobacterium tuberculosis sequence from clone y 120.	Mycobacterium leprae cosmid B971 DNA sequence.	Alcaligenes eutrophus chromsomal transketolase (cbbTc) and	phosphoglycolate phosphatase (cbbZc) genes, complete cds. Homo saciens chromosome 7 *** SEQUENCING IN PROGRESS *** 25	unordered pieces.	Homo sapiens chromosome 7, *** SEQUENCING IN PROGRESS ***, 25	unordered pieces. Homo sapiens chromosome 17, clone hRPK.372_K_20, complete sequence.	Streptomyces coelicolor cosmid 2A11		Mycobacterium tuberculosis H37Rv complete genome; segment 143/162.		Kumex acetosa expansin (EXP3) gene, partial cds.		Streptomyces lividans rpsP, trmD, rplS, sipW, sipX, sipY, sipZ, mutT genes	and 4 open reading frames.	Streptomyces coelicolor cosmid 2E1.	Streptomyces coelicolor cosmid 2E1.	Human DNA sequence from clone 173D1 on chromosome 1p36.21-	36.33.Contains ESTs, STSs and GSSs, complete sequence.	SECUENCING IN PROGRESS *** in unordered pieces	Homo sapiens chromosome X clone RP4-719K3 map q21.1-21.31, ***	SEQUENCING IN PROGRESS ***, in unordered pieces.		_	SECUENCING IN PROGRESS ***, 44 unordered pieces. Drosophila malanoaster chromosome 31 /7642 close DD0108 48815 ***	
AI 021841		AL021841	780343		¥00000	L78821	M68904	AC009541		AC009541	AC005951	AL 031184		AL021841		AF16/358		Z86111		AL023797	AL023797	AL031984	400034		AL109931			AC009367	4000367	OCEOODY.
53662	143678	53662	37085		57.104	37566	2760	169583		169583	155450	22789	155450	53662	0	1022 250445	703440	7860		38962	38962	117338	767114	107	267114		36224	226055	226055	*
GB BA1:MTV016	5	GB BA1 MTV016	5		GB_BA!:M3G1120	GB_BA1.MSGB971C	GB_BA1.AFACBBTZ	GB_HTG4_AC009541_169583		GB_HTG4.AC009541 169583	GB_PR4.AC005951	GB BA1:SC2A11	GB_PR4.AC005951	GB_BA1:MTV016		GB_PL2:AF16/358	02180005.801n_80	GB_BA2:SKZ86111		GB_BA1.SC2E1	GB_BA1:SC2E1	GB_PR2:HS173D1	ANTER ASTREAM	3	GB HTG2:HSDJ719K 267114	ı د	GB_BA1:SCD78	GB_HTG4:AC009367	CB HICA ACONO367 226065 ACON0367	10000000000000000000000000000000000000
613	:		1587	3			1296				579			591				582				1287					286			
rxa00405			7x300420				rxa00435				rxa00437			rxa00439				rxa00440				rxa00441					rxa00446			

9-Jun-98	18-OCT-	1997 18-OCT- 1997	2-Aug-99	2-Aug-99	24-Aug-99	8-Aug-97 27-Apr-99 8-Jul-99		8-Jan-98	01-MAR- 1994	17-Jun-98	17-Jun-98	01-MAR- 1994	17-Sep-98 01-MAR-	1994 03-DEC- 1999	03-DEC-	17-Feb-97
35,682	31,373	31,373	40,000	40,000	35,714	39,308 37,487 38,116		74,259	37.248	39,725	39,451	39,178	60,835 38,041	36,756	36,756	99,913
Homo sapiens	Homo sapiens	Homo sapiens	Drosophila melanogaster	Drosophila melanogaster	Homo sapiens	Mycobacterium leprae Drosophila melanogaster Trypanosoma brucei		Corynebacterium ammoniagenes	Mycobacterium leprae	Mycobacterium tuberculosis	Mycobacterium tuberculosis	Mycobacterium leprae	Streptomyces coelicolor Mycobacterium leprae	Homo sapiens	Homo sapiens	Corynebacterium glutamicum
Table 4 (continued) Home canions 12013 1 PAC RPCI1-130F5 (Roswell Park Cancer Institute	Homo Sapiens chromosome 12 clone RPCI-1 130F5 map 12q13.1, *** Homo sapiens chromosome 12 clone RPCI-1	SEQUENCING IN PROGRESS ***, 156 unordered pieces. Homo sapiens chromosome 12 clone RPCI-1 130F5 map 12q13.1, ***	SEQUENCING IN PROGRESS ***, 156 unordered pieces. Drosophila melanogaster chromosome 3 clone BACR02L16 (D715) RPCI-98 02.L.16 map 89E-90A strain y; cn bw sp. *** SEQUENCING IN PROGRESS	, 91 unordered pieces. Drosophila melanogaster chromosome 3 clone BACR02L16 (D715) RPCI-98 02.L.16 map 89E-90A strain y; cn bw sp, *** SEQUENCING IN PROGRESS	***, 91 unordered pieces. wk14a08.x1 NCI_CGAP_Lym12 Homo sapiens cDNA clone IMAGE:2412278 3's similar to gb:Y00764 UBIQUINOL-CYTOCHROME C REDUCTASE 11 KD	PROTEIN (HUMAN), mRNA sequence. Mycobacterium leprae cosmid B1779. Drosophila melanogaster cosmid clone 86E4. 927P1-2H3. TP 927P1 Trypanosoma brucei genomic clone 927P1-2H3,	genomic survey sequence.	B.ammoniagenes guaA gene.	Mycobacterium leprae cosmid B1620.	Mycobacterium tuberculosis H37Rv complete genome; segment 145/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 145/162.	Mycobacterium leprae cosmid B1620.	Streptomyces coelicolor A3(2) DNA for whiD and whiK loci. Mycobacterium leprae cosmid B1620.	Homo sapiens chromosome 6 clone RP1-225E12 map q24, ***	SEQUENCING IN PROGRESS *** in unordered pieces. Homo sapiens chromosome 6 clone RP1-225E12 map q24, ***	SEQUENCING IN PROGRESS ***, in unordered pieces. C. glutamicum lysC-alpha, lysC-beta and asd genes for aspartokinase-alpha and -beta subunits, and aspartate beta semialdehyde dehydrogenase, respectively (EC 2.7.2.4; EC 1.2.1.11).
0.003670	AE029367	AF029367	AC007824	AC007824	AI818057	298271 AL021086 AQ640325		Y10499	U00015	277165	277165	U00015	AJ010601		AL031772	X57226
34.000	148676		133361	133361	412	43254 29352 467		3866	42325	33818	33818	42325	4692	126464	126464	2803
	GB_HTG2:AE029367		_ GB_HTG2:AC007824 133361 AC007824	GB_HTG2.AC007824 133361 AC007824	GB_EST35.AI818057 412	GB_BA1.MLCB1779 GB_IN1:DMC86E4 GB_GSS15.AQ64032	ഗ	GB_BA1:BAGUAA	GB_BA2:U00015	GB_BA1:MTCY78	GB_BA1:MTCY78	GB_BA2:U00015	GB_BA1:SCAJ10601	GB_HTG2:HS225E12_126464	GB_HTG2:HS225E12_126464_AL031772	GB_BA1:CGLYS
	1143		424			975		1692			1641		, ,	6		1155
	rxa00448		rxa00450			rxa00461	rxa00465	rxa00487			rxa00488			1X400409		rxa00533

														10)3												_			
	1/-reb-9/	30-Jul-93	7-reo-97	11-Jun-93	90	66-Inf-07	10-Feb-99	24-Jun-99	26-Feb-97		21-Sep-99	17-Jun-98		28-Jan-97	6	01-DEC- 1998	24-Jun-97	17-Jun-98	5-Jun-97		09-MAR- 1995	17-Jun-98		05-DEC- 1998	08-OCT-	1997 (Rel	52, Created)	24-Jun-98	24-Jun-98	; ;
	177.66	99,391	93,856	98,701	677.00	677'06	100,000	68,003	68,185		63,187	62,401		62,205	6	86,358	62,468	60,814	66,095		64,315	64,863		98,810	98 810			98,810	99,368	<u> </u>
-	Corynebacterium glutamicum	synthetic construct	Corynebacterium glutamicum	Corynebacterium	flavescens	glutamicum	Corynebacterium	glutamicum Mycobacterium	tuberculosis Mycobacterium	tuberculosis	Streptomyces coelicolor	A3(2) Mycobacterium	tuberculosis	Mycobacterium	tuberculosis	Unknown.	Mycobacterium leprae	Mycobacterium	tuberculosis Corynebacterium	ammoniagenes	Mycobacterium leprae	Mycobacterium	tuberculosis	Unknown.	Corvnehacterium	glutamicum	•	Corynebacterium	glutamicum Corvnebacterium	glutamicum
Table 4 (continued)	t.glutamicum aspartate-semialdenyde denydrogenase gene.	ecombinant DNA fragment (Pstl-Xhol)	 glutamicum lysC-alpha, lysC-beta and asd genes for aspartokinase-alpha and -beta subunits, and aspartate beta semialdehyde dehydrogenase, rspectively (EC 2.7.2.4; EC 1.2.1.11). 	orynebacterium flavum aspartokinase (ask), and aspartate-semialdehyde	chydrogenase (asd) genes, complete cds.	una encoding previoaciendin aspanoninase.	ciglutamicum gene leuA for isopropylmalate synthase.	Nycobacterium tuberculosis H37Rv complete genome; segment 155/162.	Nycobacterium tuberculosis putative alpha-isopropyl malate synthase (leuA)	ene, complete cds.	Greptomyces coelicolor cosmid D25.	Nycobacterium tuberculosis H37Rv complete genome: segment 39/162.		Nycobacterium tuberculosis phosphoribosylformylglycinamidine synthase	(lurl.) gene, complete cds.	cequence 19 from patent US 5/26299.	Nycobacterium leprae cosmid B5.	Nycobacterium tuberculosis H37Rv complete genome; segment 36/162.	B ammoniagenes purF gene		Nycobacterium leprae cosmid B2266.	Nycobacterium tuberculosis H37Ry complete genome: segment 39/162.		Sequence 1 from patent US 5776740.	NA encoding serine hydroxymethyl transferase			DNA encoding serine hydroxymethyltransferase from Brevibacterium flavum.	DNA encoding serine hydroxymethyltransferase from Brevibacterium flavum	
	X82928	A07546	X5/226	L16848	144644	E 143 14	X70959	121125 AL022121	U88526		AL118514	Z95618		U34956	6	192052	295151	280226	X91252		U15182	Z95618		AR016483	F11273			E12594	F12594) ! !
Ş	1591	2112	2803	2957	1643	5	3492	121125	2412	!	41622	10451		2462		2115	38109	36850	1885		40123	10451		2104	2104	5		2104	2104	; ; 4
	GB_BATICGCYSCAS 1591 D	GB_PAT:A07546	GB_BAT.CGLYS	GB_BA1.CORASKD	1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	GB_PALE14314	GB_BA1.CGLEUA	GB_BA1:MTV025	GB BA1.MTU88526	t	GB_BA2:SCD25	GB BA1.MTCY7H7A	I	GB_BA1:MTU34956	4	GB_PA1:192052	GB_BA1:MLCB5	GB_BA1:MTCY369	GB BA1 BAPURF		GB_BA1:MLU15182	GB BA1:MTCY7H7A 10451		GB_PAT:AR016483	FM PAT F11273			GB_PAT.E12594	GB PAT-F12594)
		;	1386				1494				2409				4	76/			1470					1983					1425	-
			rxa00534				rxa00536				rxa00537					rxa00541			rxa00558					rxa00579					0850082	3

					Table 4 (continued)				
		GB_PAT:AR016483	2104	AR016483	Sequence 1 from patent US 5776740.	Unknown.	99,368	05-DEC- 1998	
		EM_PAT:E11273	2104	E11273	DNA encoding serine hydroxymethyl transferase.	Corynebacterium qlutamicum	99,368	08-OCT- 1997 (Ref.	
								52, Created)	
rxa00581	1092	GB_PAT:E12594	2104	E12594	DNA encoding serine hydroxymethyltransferase from Brevibacterium flavum. (Corynebacterium glutamicum	37,071	24-Jun-98	
		TAM DAT 611070	2104	E11273	ONA encoding serine hydroxymethyl transferase	Corynebacterium	37,071	08-OCT-	
			† 0 7			glutamicum		1997 (Rel.	
		GB_PAT:AR016483	2104	AR016483	Sequence 1 from patent US 5776740.	Unknown.	37,071	05-DEC-	
ARACOCA	124B	GR BATICORAHPS	2570	L07603	Copynebacterium glutamicum 3-deoxy-D-arabinoheptulosonate-7-phosphate	Corynebacterium	98,236	26-Apr-93	
1.X&UU.004	7))			glutamicum	•		
		GB_BA1:AOPCZA361 37941	37941	AJ223998	Amycolatopsis orientalis cosmid PCZA361.	Amycolatopsis orientalis	54,553	29-MAK- 1999	
			44250	***	Escherichia coli genomic DNA (16.8 - 17.1 min)	Escherichia coli	53,312	7-Feb-99	
rxa00618	1230	GB_EST19:AA802737 280	. 280	37	M06236 Sprime GM Drosophila melanogaster ovary BlueScript Drosophila	Drosophila melanogaster	39,928	25-Nov-98	
			į		melanogaster cDNA clone GM06236 Sprime, mKNA sequence.	Orosophila melanodaster	41 136	18-MAR-	T
		GB_EST28.AI534381	189	Al534381	SDU / 186. Oprime SD Drosophila melanogaster Sumerica Lt Con Control Drosophila melanogaster CDNA clone SD07186 Sprime similar to X89858: Ani FBgn0011558 PID:9927407 SPTREMBL:024240, mRNA sequence.			4 6661	J '
		MI IIMAMO IMA	4020	XAGRSA	D melanogaster mRNA for anillin protein.	Drosophila melanogaster	34,398	8-Nov-95	
rxa00619	1551	GB_BA1:MTCY369	36850	280226	Mycobacterium tuberculosis H37Rv complete genome, segment 36/162.	Mycobacterium tuberculosis	62,776	17-Jun-98	
		200 1841 005	38100	705151	Mycobacterium lenrae cosmid B5.	Mycobacterium leprae	61,831	24-Jun-97	
		GB_PAT.A60305	1845	A60305	Sequence 5 from Patent WO9708323.	unidentified	61,785	06-MAR- 1998	
rxa00620	1014	GB_PL2:AF063247	1450	AF063247	Pneumocystis carinii f. sp. ratti enolase mRNA, complete cds.	Pneumocystis carinii f. sp. ratti	41,060	5-Jan-99	
		GB_BA1:STMAPP	2069	M91546	Streptomyces lividans aminopeptidase P (PepP) gene, complete cds.	Streptomyces lividans	37,126	12-Jun-93	
		GB_HTG3:AC008763	214575	AC008763	Homo sapiens chromosome 19 clone CITB-E1_3214H19, *** SEQUENCING IN PROGRESS *** 21 unordered pieces.	nomo sapiens	40,020	ee-finy-c	
***************************************	0.10	CB INIT-CEVATES	150641	795559	Caenorhabditis elegans cosmid Y41E3, complete sequence.	Caenorhabditis elegans	36,986	2-Sep-99	
rxa000624	2	GB_EST13:AA362167 372	7 372		EST71561 Macrophage I Homo sapiens cDNA 5' end, mRNA sequence.	Homo sapiens	38,378	21-Apr-97	
		CB IMT CEVATE?	150641	795559	Caenorhabditis elegans cosmid Y41E3, complete sequence.	Caenorhabditis elegans	37,694	2-Sep-99	
rxa00626	1386	GB_BA1:MTCY369	36850		Mycobacterium tuberculosis H37Rv complete genome; segment 36/162	Mycobacterium tuberculosis	57,971	17-Jun-98	
		GR BATIMICBS	38109	795151	Mycobacterium leprae cosmid B5.	Mycobacterium leprae	58,806	24-Jun-97	
		GB_BA1:MLU15187	36138	U15187	Mycobacterium leprae cosmid L296.	Mycobacterium leprae	38,007	09-MAR- 1995	

									1	05											
6 4 0 0	66-09-1-0	29-Sep-97	29-Sep-97	3-Feb-99	29-Sep-97	4-Nov-96	17-Jun-98	3-Feb-99	27-Jan-99	5-Jul-99	21-MAY-	1993 29-Sep-97	3-Apr-98	01-DEC- 1998	17-Jun-98	17. lun 98	8	17-Jun-98	23-MAR-	6-8ng-99	6-Aug-99
97. 70	000,00	98,074	93,814	95,690	95,755	55,564	60,030	99,563	00'030	39,116	47,419	47,419	37,814	37,814	50,647	55 22B	241,00	40,300	35,750	40,634	40.634
- Access	glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium qlutamicum	Corynebacterium	Erwinia herbicola	Mycobacterium tuberculosis	Corynebacterium glutamicum	Mycobacterium bovis	Zymomonas mobilis	Unknown.	unidentified	Unknown.	Unknown	Mycobacterium	tuberculosis	tuberculosis	Mycobacterium tuberculosis	Homo sapiens	Drosophila melanogaster	Drosophila melanogaster
Table 4 (continued)	brevioacteriorii lavuiri geries for richarminoperatgoriic acid ariinotransierase and dethiobiotin synthetase, complete cds.	DNA sequence coding for desthiobiotinsynthetase.	DNA sequence coding for diamino pelargonic acid aminotransferase.	Brevibacterium flavum genes for 7,8-diaminopelargonic acid aminotransferase and dethiobiotin synthetase, complete cds.	DNA sequence coding for diamino pelargonic acid aminotransferase.	Erwinia herbicola adenosylmethionine-8-amino-7-oxononanoate transaminase Erwinia herbicola (bioA) gene, complete cds.	Mycobacterium tuberculosis H37Rv complete genome; segment 35/162.	Brevibacterium flavum gene for SecY protein (complete cds) and gene or adenylate kinase (partial cds).	Mycobacterium bovis MBE50a gene, partial cds, and MBE50b, MBE50c, preprotein translocase SecY subunit (secY), adenylate kinase (adk), methionine aminopeptidase (map), RNA polymerase ECF sigma factor (sigE50), MBE50d, and MBE50e genes, complete cds.	_	Sequence 1 from Patent US 4758514.	DNA coding of 2.5-diketogluconic acid reductase.	Sequence 9 from patent US 5693781	Sequence 9 from patent US 5726299.	Mycobacterium tuberculosis H37Rv complete genome; segment 76/162.	Mycobartarium tubarculosis H37Dy complete genome: segment 76/162	יין כסממנכיומון נסטכי כמוסטי ווסדיני כמויףוכי שליימוני, פנשיימוני, פנשיימוני מיים ביים	Mycobacterium tuberculosis H37Rv complete genome; segment 76/162.	RPCI-11-168G18.TJ RPCI-11 Homo sapiens genomic clone RPCI-11-		
8000	14003	E04041	E04040	D14083	E04040	U38519	AL021958	D14162	U77912	AF157493	100836	E00311	178753	192042	Z98268	708268	20707	Z98268	AQ420755	AC008332	AC008332
5256	7/77	675	1272	2272	1272	1290	28826	1516	7163	25454	1853	1853	1187	1187	37432	37432	1	37432	671	118545	118545
CACIA IOS.	מאַרופארפוסאס	GB_PAT:E04041	GB_PAT:E04040	GB_BA1:BRLBIOAD	GB_PAT.E04040	GB_BA2:EHU38519	GB_BA1:MTV041	GB_BA1:BRLSECY	GB_BA2:MBU77912	GB_BA2.AF157493	GB_PAT:100836	GB PAT E00311	GB_PAT:178753	GB_PAT:192042	GB_BA1:MTCI125	CB BA1:MTC1125		GB_BA1:MTCH25	GB_GSS12:AQ42075 671	GB_HTG3.AC008332 118545 AC008332	GB_HTG3.AC008332 118545 AC008332
706	Ce /			1392			999			930			1083			821	3			1035	
50000 500000	7400002			rxa00633			rxa00688			rxa00708			rxa00717			81200cm				rxa00727	

•	6-Aug-99	25-Feb-99	25-Feb-99	29-MAY- 1997	24-Jun-99	09-MAR- 1995	21-Sep-99	14-OCT- 1998	14-OCT. 1998	3-Nov-99	17-Jun-98	26-Apr-93	OB-MAR-	1999	11-MAR- 1994	28-DEC- 1998	8-Sep-99	16-Aug-93	28-Jul-99	28-Jul-99	16-Aug-93	1-Apr-93
	33,888	36,737	36,737	36,526	66,193	61,443	59,938	64,896	64,896	026,73	54,410	51,729	90£ 9£		44,308	35,571	36,044	99,539	99,539	66'236	99,885	100,000
	Drosophila melanogaster	Caenorhabditis elegans	Caenorhabditis elegans	Escheríchia coli	Mycobacterium tuberculosis	Mycobacterium leprae	Streptomyces coelicolor A3(2)	Caenorhabditis elegans	Caenorhabditis elegans	Chlamydomonas reinhardtii 57,970	Mycobacterium	tuberculosis Azotobacter chroococcum	Consthere DC 8801		Arabidopsis thaliana	Arabidopsis thaliana	Arabidopsis thaliana	1 Corynebacterium	grutamicum Corynebacterium glutamicum	grutamisam Corynebacterium		Corynebacterium glutamicum
	e 2 clone BACR48D10 (D867) RPCI-98 sp. *** SEQUENCING IN	PROGRESS***, 78 unordered pieces. Caenorhabditis elegans clone Y49F6, *** SEQUENCING IN PROGRESS ***.	2 unordered pieces. Caenorhabditis elegans clone Y49F6, *** SEQUENCING IN PROGRESS ****.	2 unordered pieces. E.coli genomic DNA, Kohara clone #319(37 4-37 8 min.).	Mycobacterium tuberculosis H37Rv complete genome; segment 40/162.	Mycobacterium leprae cosmid B2266.	Streptomyces coelicolor cosmid D25.	Caenorhabditis elegans chromosome V clone R08A5, *** SEQUENCING IN DBOCBES *** in unordered pieces.	Caenorhabditis elegans chromosome V clone R08A5, *** SEQUENCING IN	PROGRESS ***, in unordered pieces. Chlamydomonas reinhardtii putative O-acetylserine(thiol)lyase precursor	(Crcys-1A) mRNA, nuclear gene encoding organellar protein, complete cds. Mycohacterium tuberculosis H37Rv complete genome; segment 103/162.	Azotobacter chroococcum nitU. nitS, nitV, nitP, nitW, nitZ and nitM genes.	complete cds.	Cyanothece PCC 8801 NifP (nifP), nitrogenase (nifb), FdXN (tdXN), NitS (tills) Cyanothece Focosoriand Nift (nifH) gene, partial cds.	ATT 52430 AC16H Arabidopsis thaliana cDNA clone TAI306 3, mRNA	Sequence. Arabidopsis thaliana BAC F18G18 from chromosome V near 60.5 cM,	complete sequence. 701545695 A haliana, Columbia Col-0, rosette-2 Arabidopsis thaliana cDNA	cione 701943699, mktw sequence. B lactofermentum dapA and dapB genes for dihydrodipicolinate synthase and	dihydrodipicolinate reductase. gDNA encoding dihydrodipicolinate synthase (DDPS).	DNA encoding Brevibacterium dihydrodipicolinic acid synthase.	B. lactofermentum dapA and dapB genes for dihydrodipicolinate synthase and	dinydrodipicolinate reductase. C.glutamicum dapB gene for dihydrodipicolinate reductase.
		AC006789	AC006789	D90810	AL022004	U15182	AL118514	282281	282281	AF078693	783860	Mennan		AF001780	Z30506	110469 AC006258	A1998439	Z21502	E16749	E14520	Z 21502	X67737
	118545		83823	20476	68848	40123	41622	51920	51920	1492	31225	2000		6701	329	110469	455	3572	2001	2001	3572	1902
	GB_HTG3:AC008332 118545 AC008332	GB_HTG2:AC006789 83823	GB_HTG2.AC006789 83823	GB_BA1:D90810	GB_BA1:MTV043	GB_BA1:MLU15182	GB_BA2:SCD25	GB_HTG1:CER08A5	GB_HTG1:CER08A5	GR PI 2.AF078693	CONTRACTOR	GB_BAT.MILCT30	GD_DAT.AVIIVII NEO	GB_BA2:AF001780	GB_EST1:Z30506	GB_PL2:AC006258	GB_EST37:A1998439 455	GB_BA1:BLDAPAB	GB_PAT:E16749	GB_PAT.E14520	GB_BA1:BLDAPAB	GB_BA1:CGDAPB
		996			1293			1056			ć	600			1023			867			873	
		rxa00766			rxa00770			rxa00779			0	rxauu / 80			rxa00838			rxa00863			rxa00864	

28, 1:1,00		16-Aug-93	00	66-JUL-93	29-Sep-99	17-Jun-98		22-Aug-97	25-OCT-	1996	29-MAR-	1939 29-MAR-	1999	5-Feb-99	17-Jun-98	C L C	03-DEC- 1996	72- lin-99		18-Jun-98		29-MAK- 1999	29-MAR-	1999	08-OCT-	1997 (Rel. 52 Created)	10-Feb-99		29-Sep-97	29-Sep-97	29-Sep-97	10-Feb-99		29-Sep-97	
100 000		100,000	0	609,88	99,805	39,179		39,482	907'69		03,413	61 617	<u>.</u>	60,594	37,785	0	38,006	33 974	2	63,297	6	596,19	61,727		889'66		98,847		98,428	98,758	98 758	98,758		98,372	
minatoria de maria de					giutamicum Unknown.	Mycobacterium	tuberculosis	Mycobacterium leprae	, Streptomyces antibioticus	Č	streptomyces coelicolor	Strentomyces coelicolor		Pimelobacter sp.	Mycobacterium	tuberculosis	Mycobacterium	Twoanosoma huicei		Mycobacterium	tuberculosis	Streptomyces coelicolor	Streptomyces coelicolor		Conynebacterium	glutamicum	Corynebacterium	glutamicum	unidentified	Corynebacterium	glutamicum unidentified	Corynebacterium	glutamicum	Corynebacterium	giatallicalli
Table 4 (continued) NA encoding Brainbartarium dibudrodining acid conthase	described of the second of the	B. lactofermentum dapA and dapB genes for dihydrodipicolinate synthase and	dihydrodipicolinate reductase.	gunk encoding dinydrodipicolinate reductase (UDPK).	Sequence 18 from patent US 5804414.	Mycobacterium tuberculosis H37Rv complete genome; segment 122/162.		Mycobacterium leprae cosmid B22.	Streptomyces antibioticus guanosine pentaphosphate synthetase (gpsl) gene, Streptomyces antibioticus	complete cds.	Sireptomyces coelicolor A3(z), glycogen metabolism ciuster II.	Strentomyres coelicolor 43/2) alycopen metabolism clusteri		Pimelobacter sp. DNA for trehalose synthase, complete cds.	Mycobacterium tuberculosis H37Rv complete genome; segment 106/162.		Mycobacterium tuberculosis sequence from clone y222.	Sheared DNA-1014 TE Sheared DNA Transposoms british reports	Sheared DNA-1014, genomic survey sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 7/162.		Streptomyces coelicolor A3(2), glycogen metabolism cluster II.	Streptomyces coelicolor A3(2) glycogen metabolism clusterl.		DNA encoding tryptophan synthase.		Brevibacterium lactofermentum tryptophan operon.		Senomic DNA of trp operon of prepibacterium latophelmentamn.	DNA sequence of tryptophan operon.	Senomic DNA of tra opera of arealbacterium latorabeliams	Brevibacterium lactofermentum tryptophan operon.		DNA sequence of tryptophan operon.	
E14520	255	Z21502	746347	E10/37	AR038113	AL008967		Z98741	U19858	000	A3001205	A.1001205	227	D78198	281368	0	AD000010	AORSARON		Z 96071		AJ001206	AJ001205		E10963		X04960		E01688	E01375	F01688	X04960		E01375	
2001	3	3572	,	4	1411	56414		40281	2838		9 8 8	9589		2304	41230		41156	468	2	11700		9184	9589		3118		7725		7725	7726	7725	7725		7726	
CB DAT F14500		GB_BA1:BLDAPAB	1 + 4 C C C C C C C C C C C C C C C C C C	GB_PALE10/32	GB_PAT.AR038113	GB_BA1:MTV002		GB_BA1:MLCB22	GB_BA1_SAU19858		GB_BAT SCUUUTZUB	GR BA1 SCO001205		GB_BA1.D78198	GB_BA1:MTCY253		GB_BA1.MSGY222	GB GSS15/A065460 468	0	GB_BA1.MTCI418B		GB_BA1.SCO001206	GB_BA1:SC0001205		EM_PAT.E10963		GB_BA1.BLTRP		GB_PATE01688	GB_PAT:E01375	GB PAT F01688	GB BA1 BLTRP	I	GB_PAT:E01375	
		1026				059				(6/				1263					1102	;	ı			963					644				1545	
		rxa00865				rxa00867				000	rxauus/3				rxa00884					rxa00891					rxa00952					rxa00954				rxa00955	

	-66	-97		rei. rated)	66-	26-1	66 <u>.</u>		76-0	76-0	-66	76-0		10		ن	5	p-97	-66		66-d	66-	66-d	Ę	-04	}	n-98	AR-	n-98
	10-Feb-99	29-Sep-97	08-OCT-	1997 (Rel. 52, Created)	10-Feb-99	29-Sep-97	10.Feb.99) - -	29-Sep-97	29-Sep-97	10-Feb-99	29-Sep-97	70 202 05	190-67	12-5ep-93	O2-DEC.	1994	29-Sep-97	28-Jul-99		59-Sep-99	28-Jul-99	28-Sep-99	04-OCT-	1999 28. lin-99		17-Jun-98	03-MAR- 1998	17-Jun-98
	98,372	98,242	98,949		99,107	98,945	99 165	2	98,927	98,867	98,792	98,792	0 0 0 0 0 0	98,658	606,666	00 810	0.0.66	97,524	99,931	•	99,931	99,931	37,538	37,600	41.264		40,773	58,119	38,167
	Corynebacterium glutamicum	unidentified	Corynebacterium	glutamicum	Corynebacterium	glutamicum Corynebacterium	glutamicum	glutamicum	Corynebacterium	glutamicum unidentified	Corynebacterium	Corynebacterium	glutamicum	unidentified	S Corynebacterium	glutamiculii	Onkhown.	Corynebacterium	Corynebacterium	glutamicum	Unknown	I- Corynebacterium glutamicum	Gallus gallus	, Arabidopsis thaliana	confort single bidge		Mycobacterium	Streptomyces coelicolor	Mycobacterium tuberculosis
Table 4 (continued)	Brevibacterium lactofermentum tryptophan operon.	Genomic DNA of tro operon of prepibacterium latophelmentamn.	gDNA encoding tryptophan synthase.		Brevibacterium lactofermentum tryptophan operon.	Ontario and Anatomy of Street Control	seducino de la companya de la compan	Brevibacterium lactofermentum tryptophan operon.	DNA sequence of tryptophan operon.	Commis DNA of tra green of prepiparterium (atrophelmentamn	Brevibacterium lactofermentum tryptophan operon.	DNA sequence of tryptophan operon.		Genomic DNA of trp operon of prepibacterium latophelmentamn.	Corynebacterium glutamicum hom-thrB genes for homoserine dehydrogenase Corynebacterium	and homoserine kinase.	Sequence 1 from Patent WO 8809819.	DNA encoding for homoserine dehydrogenase(HDH)and homoserine	kinase(HK).	gonna encoding diaminophinerate decarboxyrase (DDO) and digmy, come synthase	Spanished 15 from patent IIS 5804414	Sequence 15 from pareit OS 3034719. DNA encoding Brevibacterium diaminopimelic acid decarboxylase and arginyl- Corynebacterium IRNA synthase.	Gallius natial mRNA for ATP-citrate lyase (ACL gene).	Genomic sequence for Arabidopsis thaliana BAC F1504 from chromosome I.	complete sequence.	Arabidopsis thaliana genome survey sequence 17 end of bAC F1407 of tor- library from strain Columbia of Arabidopsis thaliana, genomic survey	sequence. Mycobacterium tuberculosis H37Rv complete genome; segment 108/162.	S.coelicolor valS, fpgs, ndk genes.	AL021246 Mycobacterium tuberculosis H37Rv complete genome; segment 108/162.
	X04960	FO1688	E10963		X04960		E013/3	X04960	E01375	9	X04960	E01375		E01688	Y00546		109077	E01358		E16/55	A D 0 2 0 1 1 0	ARU36110 E14508	A 124564			AL087338	AL021246	Y13070	AL021246
	7725	7725			7775		97//	7725	7726	0	7725	7726		7725	3685		3685	2615	Ī	35/9	01.30	3579 3579	613			1 542	63033	3619	63033
	GB_BA1:BLTRP	OD DAT COLERB	EM PAT E10963	ı	00110.140	מס"מס"מס"מס	GB_PA1.E013/5	GB_BA1:BLTRP	GB PAT:E01375		GB_PATEUTO88 GB_BAT:BLTRP	GB PAT:E01375		GB PAT E01688	GB_BA1 CGHOMTHR 3685		GB_PAT:109077	GB_PAT:E01358		GB_PAT:E16755	0	GB_PAT:AR038110 GB_PAT:E14508	A 000 000 000 000 000 000 000 000 000 0	GB_PL2:AC007887	1	GB_GSS1.CNS00RN W	GB_BA1:MTV008	GB_BA1:SCVALSFP	GB_BA1:MTV008
			1237					1677			747				1050					1458			c i	(33			1644		
			rxa00956					×a00957			rxa00958				rxa00970					rxa00972				rxaouser			rxa00989		

2-Aug-96 26-OCT.	1999 2-Sep-99 12-Jun-98	12-Jun-98 17-OCT-	15-Jan-99 04-OCT- 1995 23-Jun-99	17-Sep-97 17-Jun-98 15-Jun-94	26-Apr-93 1-Jul-99 13-MAR- 1999	5-Aug-99 18-Apr-98 23-Jun-99	5-Aug-99 18-Apr-98 03-MAR-	29-MAR- 1999 23-Nov-99 16-Jul-97
40,841 36,416	36,416 39,172	39,172 34,661	68,275 65,935 40,454	38,636 51,989 e 38,088	53,723 34,322 36,181	99,820 75,966 38,296	100,000 65,511 52,477	43,750 37,475 37,319
, Corynebacterium glutamicum Caenorhabditis elegans	Caenorhabditis elegans Homo sapiens	Homo sapiens Homo sapiens	Streptomyces coelicolor Actinoplanes teichomycelicus Mycobacterium	Mycobacterium leprae 38,636 Mycobacterium 51,989 tuberculosis Streptococcus pneumoniae 38,088	Bacillus subtilis Homo sapiens Arabidopsis thaliana	Corynebacterium glutamicum Corynebacterium ammoniagenes Mycobacterium tuberculosis	Corynebacterium glutamicum Corynebacterium ammoniagenes Salmonella typhimurium	Limnadia lenticularis Homo sapiens Anathix ralla
Table 4 (continued) Corynebacterium glutamicum L-proline:NADP+ 5-oxidoreductase (proC) gene, Corynebacterium complete cds. glutamicum Caenorhabditis elegans chromosome IV clone Y39C12, *** SEQUENCING IN Caenorhabditis el	PROGRESS ***, in unordered pieces. Caenorhabditis elegans cosmid B0001, complete sequence. Homo sapiens clone RG038K21, *** SEQUENCING IN PROGRESS ***, 3	unordered pieces. Homo sapiens clone RG038K21, *** SEQUENCING IN PROGRESS ***, 3 unordered pieces. HS_3179_A1_G03_T7 CIT Approved Human Genomic Sperm Library D	Homo sapiens genomic clone Plate=31/9 Col=5 Kow=M, genomic survey sequence. Streptomyces coelicolor cosmid 1C2. A teichomyceticus leuC and leuD genes. Mycobacterium tuberculosis H37Rv complete genome; segment 132/162.	Mycobacterium leprae cosmid B637. Mycobacterium tuberculosis H37Rv complete genome; segment 131/162. S.pneumoniae ung gene and mutX genes encoding uracil-DNA glycosylase	Pacillus subtilis outB gene encoding a sporulation protein, complete cds. Homo sapiens clone UWGC:djs201 from 7q31, complete sequence. Arabidopsis thaliana chromosome II BAC F13K3 genomic sequence, complete sequence.	Corynebacterium glutamicum putative glutaredoxin NrdH (nrdH), NrdI (nrdI), and ribonucleotide reductase alpha-chain (nrdE) genes, complete cds. Corynebacterium ammoniagenes nrdH, nrdI, nrdE, nrdF genes. Mycobacterium tuberculosis H37Rv complete genome; segment 132/162.	Corynebacterium glutamicum putative glutaredoxin NrdH (nrdH), NrdI (nrdI), and ribonucleotide reductase alpha-chain (nrdE) genes, complete cds. Corynebacterium ammoniagenes nrdH, nrdI, nrdE, nrdF genes.	Limnadia lenticularis elongation factor 1-alpha mRNA, partial cds. Human DNA sequence from PAC 24M15 on chromosome 1. Contains tenascin-R (restrictin), EST. Anathix ralla elongation factor-1 alpha (EF-1a) gene, partial cds.
U31225 AL009026		144734 AC005052 512 AQ171808	AL031124 X84647 AL021287	Z99263 Z83018 Z21702	M15811 AC007938 AC006282	AF112535 Y09572 AL021287	AF112535 Y09572 X73226	AF063412 Z94055 U85702
1817 282838	39416 144734	144734 512	42210 2982 70287	44882 43523	1004 167237 2 92577	4363 : 6054 70287	4363	1093 134539 1240
GB_BA2.CGU31225 1817 GB_HTG1.CEY39C12_282838	GB_IN1:CEB0001 GB_HTG2:AC005052	GB_HTG2:AC005052 GB_GSS9:AQ171808	GB_BA1:SC1C2 GB_BA1:ATLEUCD GB_BA1:MTV012	GB_BA1:MLCB637 4488; GB_BA1:MTCY349 4352; GB_BA1:SPUNGMUT 1172	A GB_BA1:BACOUTB 1004 GB_PR4:AC007938 16723 GB_PL2:ATAC006282 92577	GB_BA2:AF112535 4363 GB_BA1:CANRDFGE 6054 N GB_BA1:MTV012 7028	GB_BA2:AF112535 GB_BA1:CANRDFGE N GB_BA1:STNRD	GB_IN2:AF063412 GB_PR3:HS24M15 GB_IN2:ARU85702
705	1110		1782	1131	954	2226	567	6 666
rxa00997	rxa01019		rxa01026	∝a01027	rxa01073	ra01079	жа01080	rxa01087

4004004	857	GB RA1-MTCY01B2	35938	295554	Table 4 (continued) Mycobacterium tuberculosis H37Rv complete genome; segment 72/162.	Ę	43,243	17-Jun-98
0000	3		175917	AC011632		tuberculosis Homo sapiens	36,471	19-Nov-99
		GB_HTG5:AC011632 175917 AC011632	175917		unordered pieces. Homo sapiens clone RP11-3N13, WORKING DRAFT SEQUENCE, 9	Homo sapiens	36,836	19-Nov-99
rxa01097	477	GB_BA2:AF030405	774	AF030405	unordered pieces. Corynebacterium glutamicum cyclase (hisF) gene, complete cds.	Corynebacterium	100,000	13-Nov-97
		GB_BA2:AF030405	774	AF030405	Corynebacterium glutamicum cyclase (hisF) gene, complete cds.	erium	41,206	13-Nov-97
01098	268	GB BA2:AF030405	774	AF030405	Corynebacterium glutamicum cyclase (hisF) gene, complete cds.	erium	97,933	13-Nov-97
		GB_BA1:MSGY223	42061	AD000019	Mycobacterium tuberculosis sequence from clone y223.	glutamicum Mycobacterium triberculosis	40,972	10-DEC- 1996
		CB BA1-MI CB1610	40055	Al 049913	Mycobacterium leprae cosmid B1610.	Mycobacterium leprae	61,366	27-Aug-99
rxa01100	861	GB_BA2:AF051846	738	AF051846	Corynebacterium glutamicum phosphoribosylformimino-5-amino-1- phosphoribosyl-4- imidazolecarboxamide isomerase (hisA) gene,	Corynebacterium glutamicum	97,154	12-MAK- 1998
		GB_BA2:AF060558	636	AF060558	complete cds. Corynebacterium glutamicum glutamine amidotransferase (hisH) gene.	Corynebacterium qlutamicum	95,455	29-Apr-98
		GB HTG1:HSDJ140A 221755	221755	AL109917	complete cus. Home sapiens chromosome 1 clone RP1-140A9, *** SEQUENCING IN	Homo sapiens	30,523	23-Nov-99
	<u>.</u>	9	9,5	AFOGO558	PROGRESS ***, in unordered pieces. Covnebacterium glutamicum glutamine amidotransferase (hisH) gene,	Corynebacterium	94,462	29-Apr-98
וטוווטאי	90/	GB_BAK.Ar 000000	3		complete cds.	glutamicum	0	00 174
		GB_BA1:SC4G6	36917	AL096884	Streptomyces coelicolor cosmid 4G6.	Streptomyces coelicolor A3(2)	38,378	66-INC-07
		GB_BA1:STMHISOPA 3981	A 3981	M31628	S.coelicolor histidine biosynthesis operon encoding hisD, partial cds., and	Streptomyces coelicolor	60,053	26-Apr-93
May 1104	622	GB BA1:STMHISOPA 3981	۸ 3981	M31628	hisC, hisB, hisH, and hisA genes, complete cds. S.coelicolor histidine biosynthesis operon encoding hisD, partial cds., and	Streptomyces coelicolor	58,333	26-Apr-93
			7,000	K88300 I A	hisC, hisB, hisH, and hisA genes, complete cds.	Streptomyces coelicolor	39,045	23-Jul-99
		GB_BAT:SC4G6	7-600	7500004	Management of the second of th	A3(2) Mycobacterium	60,364	24-Jun-99
		GB_BA1:MTCY336	3243/	990067	Mycdadacae Idea (2017) A Control of the Control of	tuberculosis	60 031	24. lun-99
rxa01105	1221	GB_BA1:MTCY336	32437	Z95586	Mycobacterium tuberculosis H3/Rv complete genome; segment 70/162.	tuberculosis	5,00	
		GB_BA1:MSGY223	42061	AD000019	Mycobacterium tuberculosis sequence from clone y223.	Mycobacterium tuberculosis	36,851	10-DEC- 1996
			0		Manage cosmid R1610	Mycobacterium leprae	60,902	27-Aug-99
xa01106	1449	GB_BA1:MLCB1610 GB_BA1:MSGY223	40055	AL049913 AD000019	Mycobacterium tuberculosis sequence from clone y223.	Mycobacterium tuberculosis	37,233	10-DEC- 1996

30-Jun-93	66-000-47	23-Feb-95	3-Feb-99	29-Sep-97	06-MAR-	23-Nov-99	6-Jul-98 12-Jun-98	12-Jun-98	1.Feb-99 07-OCT- 11	07-OCT- 1999	20-Nov-99	17-Jun-98	7-Jun-93	29-Nov-99	17-Jun-98	4-Aug-99 28-Aug-98	23-DEC- 1998	23-DEC- 1998	30-Nov-95
60,111	58,470	100,000	095'66	99,803	38,675	36,204	38,363 36,058	36,058	37,269 40,000	40,000	36,803	37,047	50,738	38,135	38,139	39,394 41,408	36,118	35,574	38,560
-Mycobacterium smegmatis	Mycobacterium tuberculosis	Corynebacterium	Corynebacterium	glutamicum Corynebacterium	glutamicum Aspergillus niger	Homo sapiens	Homo sapiens Homo sapiens	Homo sapiens	Triticum aestivum Homo sapiens	Homo sapiens	Arabidopsis thaliana	Mycobacterium tuberculosis	Leishmania donovani	Homo sapiens	Mycobacterium	Homo sapiens	Arabidopsis thaliana	Arabidopsis thaliana	Caenorhabditis elegans
Table 4 (continued) M.smegmatis genes hisD and hisC for histidinol dehydrogenase and histidinol-Mycobacterium smegmatis 60,111 phosphate aminotransferase, respectively	Mycobacterium tuberculosis H37Rv complete genome; segment 70/162.	Corynebacterium glutamicum acetohydroxy acid synthase (ilvB) and (ilvN)	genes, and acetohydroxy acid isomeroreductase (iIV.) gene, comprete cus. Brevibacterium flavum iIV. gene for acetohydroxy acid isomeroreductase,	complete cds. DNA encoding acetohydroxy-acid isomeroreductase.	Sequence 18 from Patent WO9706261.	Human DNA sequence from Fosmid 24E5 on chromosome 22q11.2-qter contains pavalbumin FSTs. STS.	Homo sapiens chromosome 19, cosmid F19750, complete sequence. Homo sapiens clone DJ1106H14, *** SEQUENCING IN PROGRESS ***, 42	unordered pieces. Homo sapiens clone DJ1106H14, *** SEQUENCING IN PROGRESS ***, 42	ordered icum at mo sap	IN PROGRESS ***, 31 unordered pieces. Homo sapiens chromosome 19 clone CIT-HSPC_475D23, *** SEQUENCING Homo sapiens	IN PROGRESS 31 undustrations. Arabidopsis thaliana genomic DNA, chromosome 5, P1 clone: MYH19,	complete sequence. Mycobacterium tuberculosis H37Rv complete genome; segment 47/162.	Leishmania donovani phosphoribosylpyrophosphate synthetase gene,	complete cds. Homo sapiens chromosome 1 clone RP4-799D16 map p34.3-36.1, *** SEQUENCING IN PROGRESS ***, in unordered pieces.	Mycobacterium tuberculosis H37Rv complete genome, segment 69/162.	Homo sapiens mRNA for KIAA1109 protein, partial cds. HS_3098_A1_C03_T7 CIT Approved Human Genomic Sperm Library D Homo sapiens genomic clone Plate=3098 Col=5 Row=E, genomic survey	sequence. Arabidopsis thaliana chromosome 1 BAC F508 sequence, complete	sequence. Arabidopsis thaliana chromosome 1 BAC F5O8 sequence, complete	sequence. Caenorhabditis elegans cosmid C06G1.
X655/2	929	L092 2	D14561	E082 2	A602 9	21.5	AC005265 AC004965	AC001965	U558 9 AC01 469	AC01 469	AB01 077	2925:9	M76513	AL05 344	274020	AB02 032 AQ10 201	AC00 990	AC00 990	U410'4
	137 2955					35506 2821	43900 AC 323792 AC		36	3436 AC	77380 AE	38970 Z9	1887 M	0149 AI	35377 Zi		99923 A	99923 A	31205 U
GB_BA1:MSHISCD 2298	GB_BA1:MTCY336 32437	GB_BA1:CORAIA 4705	GB_BA1:BRLILVCA 1364	GB PAT:E08232 1017	GB_PAT_A60299 2869	GB_PR3:HS24E5 355	GB_PR3:AC005265 436 GB_HTG2:AC004965_323		GB_PL2:TAU55859 2397 GB_HTG3:AC011469 11343	GB_HTG3:AC011469 113436	GB_PL1:AB010077 77	GB_BA1:MTCY10G2 38	GB_IN1:LEIPRPP 18	GB_HTG2:HSJ799D1 130149	GB_BA1:MTCY48 35	GB_PR2:AB029032 637: GB_GSS9:AQ107201 355	GB_PL2:F508 99	GB_PL2.F5O8 99	GB_IN1:CELC06G1 31
		1137			1449		846		1528			1098			2556		873		
		rxa01145			rxa01162		rxa01208		rxa01209			rxa01215			rxa01239		rxa01253		

05-MAY-	1999	2-Aug-99	26-OCT- 1999	15-OCT- 1998	12-Apr-99	01-OCT- 1999	11-Jun-99	23-Nov-99	6-Jul-9	28-Sep-99 ^r 9-Jul-98		23-Nov-97 20-Nov-99	-	28-Jul-99	24-Jun-99	24-Feb-97	23	91016 91016	24-Feb-97	28-Jul-99	
41,121	<u>.</u>	40,634	38,290	34,311	34,311	37,722	38,492	39,738	46,237	45,574 44,097		41,316 36,606	!	37,916	37,419	34,831		35,138 37,277	100,000	38,400	
Acedices CHOL	one saprens	Drosophila melanogaster	Drosophila melanogaster	Arabidopsis thaliana	Arabidopsis thaliana	Homo sapiens	Gossypium hirsutum	Homo sapiens	Mus musculus	Mus musculus Mus musculus		Mus musculus Arabidopsis thaliana		Homo sapiens	Mycobacterium	tuberculosis Corynebacterium	glutamicum	Streptomyces coelicolor Homo sapiens	Corynebacterium	giutarilicuiri Homo sapiens	
		chromosome 2 clone BACR38D12 (D590) RPCI-98 rain y, cn bw sp. *** SEQUENCING IN PROGRESS	•••, 60 unordered pieces. Drosophila melanogaster chromosome 2 clone BACR35F01 (D1156) RPCI-98 Drosophila melanogaster. 35.F.1 map 48A-48C strain y; cn bw sp, ••• SEQUENCING IN PROGRESS	••• 108 unordered pieces. Arabidopsis thaliana chromosome II BAC F12A24 genomic sequence,	ı chromosome II BAC T24121 genomic sequence,	complete sequence. Homo sapiens clone 4_K_17, LOW-PASS SEQUENCE SAMPLING.	BNLGHi12371 Six-day Cotton fiber Gossypium hirsutum cDNA 5' similar to	Human DNA sequence from PAC 227P17, between markers DXS6791	and UX 58038 on chromosome A contains of contains AV171099 Mus musculus head C57BU6J 14, 17 day embryo Mus musculus	cDNA clone 3200002M11, mRNA sequence. Mus musculus mGpi1 gene, exon 1	uc83d10.y1 Sugano mouse kidney mkia Mus muscuus cura cione IMAGE:1432243 5' similar to TR:035120 035120 MGPI1P. ;; mRNA	sequence. Mus musculus mRNA for mGpi1p, complete cds. Mus musculus mRNA for mGpi1p, complete cds. P1 clone: MJJ3, complete Arabidopsis thallana	Arabidopsis thallana genomic DINA, cilifornosonic 3, 17 construction	sequence. HS_2026_A2_C09_T7C CIT Approved Human Genomic Sperm Library D Homo sapiens genomic clone Plate=2026 Col=18 Row=E, genomic survey	sequence. Maccharterium tuberculosis H37Rv complete genome; segment 40/162.		C glutamicum lyse and lyse genes.	Streptomyces coelicolor cosmid 5A7. Homo sapiens chromosome 4 clone B220G8 map 4q21, complete sequence.	C.glutamicum lysE and lysG genes.		
	AQ518843	AC007473		AC005167	AC005825		AI725583	Z81007	AV171099	AB008915	AI050532	AB008895	AB005237	AQ766840	*00000		X96471	AL031107 4 AC004054	X96471		AQ769223
	441	194859	115847	83260	97380	127222	728	82951	9 173	530	293	3062	87835	1 491	9	08848	2374	40337			3 500
	GB_GSS14;AQ51884 441	3 GB_HTG2.AC007473 194859	GB_HTG4.AC011696 115847 AC011696	GB PI 2.ATAC005167 83260	GB PI 2 ATAC005825 97380	GB_HTG3:AC011150 127222	GB_EST32:AI725583	GB_PR2:HS227P17	GR EST34-AV171099 173	GB RO:AB008915S1	GB_EST22:AI050532	GB_RO:AB008895	GB_PL1:AB005237	GB_GSS5:AQ766840_491	3	GB_BA1:MTV043	GB_BA1:CGLYSEG	GB_BA1:SC5A7	030000000000000000000000000000000000000	66_6A!.CGC! 3CC	GB_GSS5:AQ769223 500
	1044			706	3		259			629	3		944				993			778	
	rxa01321			04040	1X401332		rxa01360			~ 201361	200		rxa01381				xa01393			xa01394	

]	14																		_
	17-Jun-98	18- Jun-98		27-Aug-99	20-OCT- 1995	17-Apr-96	17-Apr-96	24-Jun-99	13-Aug-99	20-Aug-99	3-Jun-99	19-Aug-99	13-Apr-99	7-Jan-99	26-Apr-93	20-Aug-97	20-Aug-97	29-Jun-99	01-OCT-	1998	01-OCT- 1998	17-Jun-98	!	17-DEC- 1993	8-Jul-99	2-Sep-99	07-DEC-	1999	27-Apr-93	17-Jun-98	29-Sep-94
	40.086	43 343)))	38,177	64,876	38,943	37,500	38,010	36,346	37,897	36 149	35,846	40,566	38,095	38,206	36,623	34,719	37,500	37,031		38,035	38,371		38,064	60,775	38,514	37,730		39,340	63,300	36,756
	Mycobacterium	tuberculosis	Mycobacterium tuberculosis	Mycobacterium leprae	Euglena gracilis	Escherichia coli	Escherichia coli	Mycobacterium tuberculosis	Drosophila melanogaster	Drosophila melanogaster	Total concept of the second	Drosophina meranogaster Arabidopsis thaliana	Sorosporium saponariae	Arabidopsis thaliana	Anabaena sp.	Homo sapiens	Homo sapiens	Mus musculus	Homo sapiens		Homo sapiens	Mycobacterium	tuberculosis	Escherichia coli	Streptomyces coelicolor	Caenorhabditis elegans	Homo sapiens		Sus scrofa	Mycobacterium	Mycobacterium leprae
Table 4 (continued)	Mycobacterium tuberculosis H37Rv complete genome; segment 154/162.	Wilder of the state of the stat	Mycobacterium tuberculosis H37Rv complete genome; segment 153/162.		Mycobacterium lepide Cosmin Lebras. E.gracilis mRNA for GTP cyclohydrolase I (core region).	solution 1 00 ct 8 50 most accional	Escherichia coli K-12 chromosomal region non 32.8 to 00.1 minutes	Escherichia coli K-12 chromosomal region from 32.3 to 00.1 militatos. Mycobacterium tuberculosis H37Rv complete genome; segment 93/162.	Drosophila melanogaster mRNA for drosophila dodeca-satellite protein 1	(DDP-1). Drosophila melanogaster chromosome 2 clone BACR01106 (D1054) RPCI-98	***, 86 unordered pieces.	Drosophila melanogaster clone LD21677 unknown mRNA.	Arabidopsis thaliana BAC F6H8. Sorosporium saponariae internal transcribed spacer 1, 5.8S ribosomal RNA.	gene; and internal transcribed spacer 2, complete sequence. Arabidopsis thaliana chromosome II BAC T15J14 genomic sequence.	complete sequence. Anabaena sp. (clone AnH20.1) nitrogen fixation operon nifB, fdxN, nifS, nifU,	and nifH genes, complete cds.	Human BAC clone RG204116 from 7431, complete sequence.					sequence.	Mycobacterium (uperculosis flority complete general cognition	E. coli chromosomal region from 89.2 to 92.8 minutes.			Caenornabditis elegans cosmid 102197, comprete cognitive and contract and X-2 (DI X-2) dene, complete cds.		Dia Diamino acid oxidase (DAO) gene, exon 1.	Mycobacterium tuberculosis H37Rv complete genome; segment 76/162.	Mycobacterium leprae cosmid L247.
	705/36		295557		AL023093 Z49757		U14003	U14003 Z73966	A.1238847	AC009210		AF132179	AF178045 AF038831	AC005957	J05111		AC002461	AC002461	AL049866	AC005740	AC005740		284724	00000		AL096823	AL032630	03100	****	Z98268	U00021
	. 03060		24244		38916 242			338534 39430	5419			4842	82596	7 108355	5936		197273	197273	165901	186780	186780		0 35420	176195				3202		395 37432	39193
		GB_BA1:MIY15C1U	GB_BA1:MTCY7H7B		GB_BA1:MLCB2548 GB_PL1:EGGTPCHI		GB BA1:ECOUW93	GB_BA1:ECOUW93 GB_BA1:MTCY49	CB 1814-DMRE238847	GB_HTG3:AC009210 103814		GB IN2:AF132179	GB_PL2:F6H8	GB_FLZ.AFU38831	GB BA1.ANANIFBH		GB PR2:AC002461	GB_PR2:AC002461	GB_RO:MM437P9	GB_PR3:AC005740	GB PR3:AC005740		GB_BA1:MTCY22G10 35420	GB BA2-ECOUW89		GB_BA1:SCQ11	GB_IN1:CEY62H9A	GB_PR4:HSU51003		GB_OM:PIGDA01	GB BA1:U00021
			711				975			2			009		021	176			651				1998				1053			1785	3
			rxa01514 7				rxa01515 9			olcioaxi			rxa01517		2001521				cca01528				rxa01551				rxa01561			0011000	000

	70 211 62	5-Jul-99		23-Nov-99	6-Jul-99		17-Jun-98	9-Feb-96	7 MAY	1999	17-Jun-98	30 0110 05	20-50-03	05-MAT- 1999	28-OCT-	1997	28-OCT- 1997	28-OCT- 1997	100	C6-40N-67	14-Jun-96	1-Jun-99	15-Jan-99	24-Nov-98	2-Jul-99	3-Jun-98		27-Apr-93	14-Jul-95	04-MAY-	998 1	17-Jun-98
	247 20	30,730 40,811		38,768	39,018		40,656	44,262	7	40,703	40,986	25 254	100.00	35,354	41,894		41,712	39,576		39,157	39,157	38,910	60,644	38,037	36.122	48,079		37,093	37,093	100,000		36,323
-		Mycobacterium leprae Homo sapiens		Homo sapiens	Homo sapiens		Mycobacterium tuberculosis	Homo sapiens		I niobaciilus remooxidans	Mycobacterium	tuberculosis Mais mitoribis	אומס ווומסכחומס	Mus musculus	Tula virus		Tula virus	Tula virus		Homo sapiens	Homo sapiens	Gossypium robinsonii	Streptomyces coelicolor	Drosophila melanogaster	Drosophila melanogaster			Rattus norvegicus	Rattus sp.	Corynebacterium	glutamicum	Mycobacterium tuberculosis
	Table 4 (continued)	Mycobacterium leprae cosmid B1351. Human chromosome Xq28, cosmid clones 7H3, 14D7, C1230, 11E7, F1096,	A12197, 12G8, A09100, complete sequence bases 1, 217657.	Homo sapiens DNA sequence from PAC 13D10 on chromosome 6p22.3-23.	Contains CpG island. Human chromosome Xq28, cosmid clones 7H3, 14D7, C1230, 11E7, F1096.	A12197, 12G8, A09100, complete sequence bases 1217657.	Mycobacterium tuberculosis H37Rv complete genome; segment 117/162.	HUM213D06B Human aorta polyA+ (TFujiwara) Homo sapiens cDNA clone	GEN-213D06 5', mRNA sequence.	Thiobacillus ferrooxidans carboxysome operon, complete cds.	Mycobacterium tuberculosis H37Rv complete genome; segment 134/162.		M. musculus retrovirus restriction gene nv i.	Sequence 1 from Patent WO9743410.	Tula virus 064 nucleocapsid protein gene, partial cds.		Tula virus O52 nucleocapsid protein gene, partial cds.	Tula virus O24 nucleocapsid protein gene, partial cds.		ys81e01.s1 Soares retina N2b4HR Homo sapiens cDNA clone IMAGE:221208 3' similar to gb:X63749_rna1 GUANINE NUCLEOTIDE- BINDING PROTEIN G(T), ALPHA-1 (HUMAN);, mRNA sequence.	human STS SHGC-30023, sequence tagged site.	Gossypium robinsonii CelA2 pseudogene, partial sequence	Streptomyces coelicolor cosmid 1C2.	GH04563.5prime GH Drosophila melanogaster head pOT2 Drosophila	Increasely asset CDIVA cione of 104303 opinios, mixes sequence. Decembile melanoraster periodentide E (not) gene complete cds	Lactobacillus reuteri cobalamin biosynthesis protein J (cbiJ) gene, partial cds;	and uroporphyrin-III C-methyltransferase (sumT) gene, complete cds.	Rat heavy neurofilament (NF-H) polypeptide, partial cds.	Rat mRNA for heavy neurofilament polypeptide NF-H C-terminus.	Corynebacterium glutamicum chorismate synthase (aroC), shikimate kinase	(aroK), and 3-dehydroquinate synthase (aroB) genes, complete cds; and putative cytoplasmic peptidase (pepQ) gene, partial cds.	Mycobacterium tuberculosis H37Rv complete genome; segment 111/162.
	!	Z95117 AL034384		AL021407	AL034384		Z95387	D79278		AF129925	AL021309		81//6 X	A67508	U95309		U95303	U95302		H91843	G26925	AF139451	AL031124	A1064232	A E 117806	AF067123		M37227	X13804	AF124600		Z83863
		38936 217657		153147	217657		25949	392		10243	11364		6480	6480	900		900	900		362	362	1202	42210		000	1034		3085	3085	4115		33818
		GB_BA1:MLCB1351 GB_PR2:HSMTM0		GB_PR2:HS13D10	GB PR2 HSMTM0) 	GB_BA1:MTCY1A10	GB_EST6:D79278		GB_BA2:AF129925	GB_BA1:MTV013		GB_RO:MMFV1	GB_PAT:A67508	GB_VI:TVU95309		GB_VI:TVU95303	GB_VI:TVU95302		GB_EST5:H91843	GB STS G26925	GB PL2:AF139451	GB_BA1:SC1C2	GB_EST22.AI064232	0001417.0141.000	GB_BA2.AF117830	ı	GB_RO:RATNFHPEP	GB_RO:RSNFH	GB_BA2:AF124600		GB_BA1.MTCY159
		795	2				723				675				651					1359			1224			873				1353		
		rva01617	0				rxa01657				xa01660				xa01678					xa01679			Cx a0 1690			xa01692				rxa01698		

15-Jun-96	04-MAY- 1999	07-DEC- 1997	24-OCT- 1996	29-OCT-	29-OCT- 1998	23-Nov-99	23-Nov-99	09-MAR- 1999	18-MAR- 1999	12-Jul-97	18-MAK- 1999	22-MAY- 1995	22-MAY- 1995	17-Jun-98	22-Aug-97	22-Jul-99	23-Jun-98	2-Apr-98	12-Apr-98
62,780	100,000	40,260	45,425	40,876	41,367	35,651	35,651	39,671	35,817	35,698	37,243	42,812	42,655	59,294	57,584	61,810	39,655	35,942	40,000
Mycobacterium leprae	Corynebacterium glutamicum	Streptomyces caelestis	Oryza sativa	i Trypanosoma cruzi	ti Trypanosoma cruzi	Homo sapiens	Homo sapiens	Mus musculus	Homo sapiens	Arabidopsis thaliana	Homo sapiens	Homo sapiens	Homo sapiens	Mycobacterium tuberculosis	Mycobacterium leprae	Streptomyces coelicolor A3(2)	Homo sapiens	Rattus sp.	IA Neurospora crassa
Table 4 (continued) Mycobacterium leprae cosmid B937 DNA sequence.	Corynebacterium glutamicum chorismate synthase (aroC), shikimate kinase (aroK), and 3-dehydroquinate synthase (aroB) genes, complete cds; and	putative cytoplasmic peptidase (pepul) gene, partial cus. Streptomyces caelestis cytochrome P-450 hydroxylase homolog (nidi) gene, partial cds; polyketide synthase modules 1 through 7 (nidA) genes, complete	cds; and N-methyltransferase homolog gene, partial cds. C19712 Rice panicle at ripening stage Oryza sativa	mRNA sequence. TENS1404 T. cruzi epimastigote normalized cDNA Library Trypanosoma cruzi Trypanosoma cruzi	cDNA clone 1404 5', mRNA sequence. TENS1404 T. cruzi epimastigote normalized cDNA Library Trypanosoma cruzi Trypanosoma cruzi cDNA clone 1404 5', mRNA sequence.	Homo sapiens chromosome 1 clone RP4-534K7, *** SEQUENCING IN	PROGRESS ***, in unordered preces. Homo sapiens chromosome 1 clone RP4-534K7, *** SEQUENCING IN	PROGRESS ***, in unordered pieces, mg9191608 x 1 Stratagene mnouse heart (#937316) Mus musculus cDNA clone mg91016101 musculus cDNA clone	IMACE: 300 110 3, minuty sequence. Homo sapiens PAC clone DJ1060B11 from 7q11.23-q21.1, complete	sequence. Arabidoosis thaliana BAC TM018A10.	Homo sapiens PAC clone DJ1060B11 from 7q11.23-q21.1, complete	sequence. yas203.s.1 Soares infant brain 1NIB Homo sapiens cDNA clone yas707.32000.01 mDNA camience	yg52a03.s1 Soares infant brain 1NIB Homo sapiens cDNA clone yg62a00.s1, mRNA sequence.	Mycobacterium tuberculosis H37Rv complete genome: segment 98/162.		Mycobacterium leptae cosmiu 522. Streptomyces coelicolor cosmid 5F7.	om38c02.s1 Soares_NFL_T_GBC_S1 Homo sapiens cDNA clone IMAGE:1543298 3' similar to WP.F28F8.3 CE09757 SMALL NUCLEAR	RIBONUCLEOPROTEIN E.; mRNA sequence. EST110563 Rat PC-12 cells, NGF-treated (9 days) Rattus sp. cDNA clone	RPNBI81 5' end, mRNA sequence. NCP6G8T7 Perithecial Neurospora crassa cDNA clone NP6G8 3' end, mRNA Neurospora crassa sequence.
L78820	AF124600	AF016585	C19712	AA952466	AA952466	AL109925	AL109925	AI447108	179640 AC006322	AF013294		R46227	R46227	270283	!	298741 AL096872	AA918454	H34042	AA899038
38914	4115	41097	399	5 278	5 278	< 154416	< 154416	431	179640	106184	179640	443	443	34150		40281	54 416	345	38 450
GB_BA1:MSGB937C	S GB_BA2:AF124600	GB_BA2:AF016585	GB_EST9.C19712	_ GB_EST21:AA952466 278	_ GB_EST21:AA952466 278	GB_HTG1:HSDJ534K 154416 AL109925	7 GB_HTG1:HSDJ534K 154416	7 GB_EST27.AI447108	GB_PR4:AC006322	CB BISTMU18410	GB_PR4:AC006322	GB_EST3.R46227	GB_EST3:R46227	GB_BA1:MTCY190		GB_BA1:MLCB22 GB_BA1:SC5F7	GB_EST21:AA918454 416	GB EST4:H34042	_ GB_EST20:AA899038 450
	693			805		684			1332			876		1167			924		
	rxa01699			xa01712		xa01719			rxa01720			rxa01746		rxa01747			α a 01757		

												11	7																
	22-Jun-99 16-OCT-	1999	16-0CT- 1999	7-Jan-99	15-DEC-	1994	15-Feb-99	26-Apr-99	26-MAR- 1999	8-Apr-99	U18997	12-Nov-98	29-MAY-	1888	14-OCT- 1998	14-OCT.	1998	14-0CT-	1998	19-Jun-98	27-Aug-99	15-Jun-96	19-Jun-98	29-Nov-96	18-Aug-99	5-Aug-99	-	18-Apr-98	
	40,067 35,450		35,450	100,000	38,692		36,962	38,109	37,021	37,021	37,196	38,021	39,860		37,564	37 564	97,364	37,576		35,910	64,260	64,260	37,229	38,525	31,579	99,733		70,321	
	Aeropyrum pernix Drosophila melanogaster		Drosophila melanogaster	Corynebacterium	glutamicum Rattus norvegicus		Anas platyrhynchos	Zea mays	Homo sapiens	Homo sapiens	Escherichia coli	Escherichía coli	Haemophilus influenzae		Caenorhabditis elegans	and the state of t	Caenornabuilis elegaris	Caenorhabditis elegans	•	Mycobacterium tuberculosis	Mycobacterium leprae	Mycobacterium leprae	Mycobacterium tuberculosis	Caenorhabditis elegans	8 Drosophila melanogaster	Corynebacterium	glutamicum	Corynebacterium ammoniagenes	
Table 4 (continued)	Aeropyrum pernix genomic DNA, section 6/7. Drosophila melanogaster clone RPC198-6H2, *** SEQUENCING IN	PROGRESS ***, 75 unordered pieces.	Drosophila melanogaster clone RPC198-6H2, *** SEQUENCING IN DROGRESS *** 75 inordered pieces	Corynebacterium glutamicum 3' ppc gene, secG gene, amt gene, ocd gene	land 5' soxA gene. Battus nonvenicus (clone A21 I42) alnha2u diphulin gene exons 1-7.	ייאמונים וייניסקיים (מוסיס לייסיס (מיסיס איניס מיסיס	Anas platyrhynchos (Super M) IgY upsilon heavy chain gene, exon 2.	486101D10 x1 486 - leaf primordia cDNA library from Hake lab Zea mays	cDNA, mRNA sequence. SHGC-62915 Human Homo sapiens STS genomic, sequence tagged site.	RPC11-4112.TV RPCI-11 Homo sapiens genomic clone RPCI-11-4112,	genomic survey sequence. Escherichia coli K-12 chromosomal region from 67.4 to 76.0 minutes.	Escherichia coli K-12 MG1655 section 282 of 400 of the complete genome.	_		Caenorhabditis elegans chromosome IV clone Y64F11, *** SEQUENCING IN	PROGRESS, in unordered pieces.	Caenorhabditis elegans chromosome IV clone Y64-11, *** SEQUENCING IN PROGRESS *** in unordered pieces.	Caenorhabditis elegans chromosome IV clone Y64F11, *** SEQUENCING IN Caenorhabditis elegans	PROGRESS ***, in unordered pieces.	Mycobacterium tuberculosis H37Rv complete genome; segment 126/162.	Mycobacterium leprae cosmid B250.	Mycobacterium leprae cosmid B1529 DNA sequence.	Mycobacterium tuberculosis H37Rv complete genome, segment 126/162.	Caenorhabditis elegans cosmid F46H5.	Drosophila melanogaster chromosome 2 clone BACR03E19 (D1033) RPCI-98 Drosophila melanogaster 03.E.19 map 36E-37C strain y, cn bw sp, *** SEQUENCING IN PROGRESS	, 34 unordered pieces. Corvinebacterium alutamicum ribonucleotide reductase beta-chain (nrdF)		Corynebacterium ammoniagenes nrdH, nrdI, nrdE, nrdF genes.	
	185300 AP000063		AC010694	AJ007732	M24108	7100	X78272	AI629479	G48245	B49052	110000 U18997	AE000392	U32715		299776		299776	799776		274024	Z97369	L78824	274024	1141543	AC009204	AF112536		Y09572	
	185300		115857	4460	7501	200	1381	353	515	515	110000	10345	13136		177748	,	177748	177748		39991	40603	36985	39991	3888		1798	3	6054	
	GB_BA1:AP000063	- - - - - - - - -	GB_HTG4:AC010694	GB_BA1:CGL007732	0.00	פם_אט.האוארפר	GB OV:APIGY2	GB_EST30:AI629479	GB_STS.G48245	GB_GSS3:B49052	GB_BA2:ECOUW67_	0 GB_BA2:AE000392	GB_BA2:U32715		GB_HTG1;CEY64F11 177748		GB_HTG1:CEY64F11	GB HTG1:CEVE4E11 177748		GB_BA1:MTCY274	GB BAT MLCB250	GB_BA1:MSGB1529C 36985	GB_BA1:MTCY274	CB INT-CELEAGHS	GB_HTG3:AC009204	GB BA2.4F112536		GB_BA1:CANRDFGE 6054 N	
	915			401				654			1470				1002					852			978			1125	7		
	rxa01807			rxa01821				rxa01835			rxa01850				rxa01878					rxa01892			rxa01894			0201057	0.26 0.64		

												1	.18												
	23-Apr-98	11-MAY-	1999	15-Sep-99	15-Sep-99	11-MAY-	1999	19-Sep-96	14-Sep-93	18-Jun-96	20-Jan-99	26-Sep-99 8-Aug-95	G.	30-Nov-97	17-Feb-96	17lun-98		14-Jun-96	09-MAR- 1995			24-MAY- 1993	30-Jul-99	6-Feb-99	
	72,082	100,000		35,917	33,925	100,000		38,749	39,305	61,417	38,560	40,275 100 000	2	38,889	36,647	59.415	7	57,093	57,210			99,317	94,387	sa 62,247	
	Corynebacterium	ammoniagenes Coxnebacterium	glutamicum	Chloroplast Arabidopsis thaliana	Chloroplast Arabidopsis	Corynebacterium	glutamicum	Xanthomonas campestris pv. vesicatoria	Xanthomonas campestris	Crithidia fasciculata	Helicobacter pylori J99	Mus musculus	glutamicum	Corynebacterium	Anabaena PCC7120	M. Control of the Con	tuberculosis	Mycobacterium leprae	Mycobacterium leprae			Corynebacterium	Corynebacterium	giutamicum Pseudomonas aeruginosa 62,247	
Table 4 (continued)	Convnebacterium ammoniagenes ribonucleoside diphosphate reductase small Corynebacterium	subunit (nrdF) gene, complete cds.	C glutamicum panB, panC & xylb genes.	Arabidopsis thaliana chloroplast genomic DNA, complete sequence.	strain:Columbia. Arabidopsis thaliana chloroplast genomic DNA, complete sequence,	strain:Columbia.	C.giulalliculii pailu, pailu, pailu, pailu	Xanthomonas campestris hrpB pathogenicity locus proteins HrpB1, HrpB2, HrpB3, HrpB4, HrpB5, HrpB6, HrpB7, HrpB8, HrpA1, and ORF62	genes, complete cds. Xanthomonas campestris hrpB6 gene, complete cds.	Crithidia fasciculata inosine-uridine preferring nucleoside hydrolase (IUNH)	gene, complete cds. Helicobacter pylori, strain J99 section 28 of 132 of the complete genome.	Homo sapiens Leman coiled-coil protein (LCCP) mRNA, complete cds.	C.glutamicum dapE gene and orf2.	C.glutamicum ORF3 and aroP gene.	Anabaena PCC7120 nitrogen fixation proteins (niff., nifN, nifX, nifW) genes.	complete cds, and nitrogenase (nifk) and hesA genes, partial cds.	Mycobacterium tuberculosis H37Rv complete genome, segment 52/162.	M. leprae genomic dna sequence, cosmid b1912.	Mycobacterium leprae cosmid B1756.			C.glutamicum GDHA gene.	Corynebacterium glutamicum, gdh gen for glutamate dehydrogenase.	Pseudomonas aeruginosa gdhA gene, strain PAC1.	
	AE050168		X96580	AP000423	AP000423		X3658U	U33548	M99174	U43371	AE001467	AF175967	X81379	X85965	1147055		293777	L01536	U15180			X72855	X59404	Y18494)
	122B	077	2164	154478	154478		2164	8429	1329	1060	11601	3492	1966	2612	6460	50	29540	38503	38675			2037	2037	1628	2
	000000000000000000000000000000000000000	GB_BAZ.Arusu108	GB_BA1:CGPAN	GB_PL1:AP000423	GB PL1:AP000423	1	GB_BA1:CGPAN	GB_BA1.XCU33548	GB_BA1;XANHRPB6 1329	A GB_IN2 7F143371	74467	GB_BAZ 31467 GB_RO Al : 75967	GB_BA1.CGDAPE	GB BA1:CGDNAARO 2612	q 9	GB_BATAPU47033	GB_BA1:MTCI364	GB_BA1:MSGB1912C 38503	S GB_BA1:MLU15180			GB BA1:CGGDHA	GB BA1:CGGDH	CD_D/11:00000000000000000000000000000000000	GB_BAI.FAE 1017
			096				936			1059			1230				859					1464			
			rxa01928				xa01929			rxa01940			rxa02022				rxa02024			xa02027	rxa02031	2702022	1		

17-Jun-98	24-Jun-97	29-MAY- 1995	4-Jun-97	27-0CT-	1997 6-Nov-97	13-Jan-99	31-DEC- 1998	18-MAY-	1995	24-MAR- 1999	01-MAR- 1994	24-Jun-99	24-Sep-99	14-MAY-	24-Sep-99	02-MAR- 1998	11-Jun-99	02-MAR-	23-Nov-99	9-Jun-99	26-Jun-98
38,442	56,486	52,127	34,163	35,586	31,917	35,818	34,274	41,162		50,791	37,563	39,504	37,909	37,843	37,909	36,533	33,451	36,756	34,365	34,325	33,874
Mycobacterium	Mycobacterium leprae	Escherichia coli	Homo sapiens	Homo sapiens	Homo sapiens	Streptomyces coelicolor	Homo sapiens	Homo sapiens		Streptomyces coelicolor	Mycobacterium leprae	Mycobacterium tuberculosis	8 Drosophila melanogaster	Arabidopsis thaliana	8 Drosophila melanogaster	Streptomyces coelicolor	Gossypium hirsutum	Streptomyces coelicolor	Homo sapiens	Arabidopsis thaliana	Arabidopsis thaliana
Table 4 (continued) Mycobacterium tuberculosis H37Rv complete genome; segment 49/162.	Mycobacterium leprae cosmid B33.	E. coli genomic sequence of the region from 84.5 to 86.5 minutes.	zw82h01.r1 Soares_testis_NHT Homo sapiens cDNA clone IMAGE:782737	no 18810.11 NCI_CGAP_GCB1 Homo sapiens cDNA clone IMAGE:1183963	5", mKNA sequence. Himan PAC clone DJ0596009 from 7o15, complete sequence.	Streptomyces coelicolor cosmid 1A6.	Homo sapiens chromosome 17, clone hRPK 112_J_9, complete sequence.	yg71g10.r1 Soares infant brain 1NIB Homo sapiens cDNA clone	IMAGE:38768 5' similar to gb:V00567 BETA-2-MICROGLOBULIN PRECURSOR (HUMAN);, mRNA sequence.	Streptomyces coelicolor cosmid 6G10.	Mycobacterium leprae cosmid B1170.	Mycobacterium tuberculosis H37Rv complete genome; segment 70/162.	Drosophila melanogaster chromosome 3 clone BACR09D08 (D1101) RPCI-98 Drosophila melanogaster 09.D.8 map 96F-96F strain y; cn bw sp, *** SEQUENCING IN PROGRESS	T12A12-Sp6 TAMU Arabidopsis thaliana genomic clone T12A12, genomic	Suivey sequence. Drosophila melanogaster chromosome 3 clone BACR09D08 (D1101) RPCI-98 Drosophila melanogaster 09.D.8 map 96F-96F strain y; cn bw sp, *** SEQUENCING IN PROGRESS	•••, 121 unordered pieces. S.coelicolor secY locus DNA.	BNLGHi10185 Six-day Cotton fiber Gossypium hirsutum cDNA 5' similar to (AC004005) putative ribosomal protein L7 [Arabidopsis thaliana], mRNA	sequence. S.coelicolor secY locus DNA.	Human DNA sequence from clone RP3-525L6 on chromosome 6p22.3-23	Contains CA repeat, 313s, GSSS and a Cho Island, Complete Sequence. Arabidopsis thaliana DNA chromosome 4, BAC clone F21P8 (ESSA project).	Arabidopsis thaliana BAC T7123, complete sequence.
295585	294723	M87049	AA448146	AA641937	AC003074			R49746		AL049497	U00010	Z95586	AC010579	B09839	AC010579	X83011	AI731596	X83011	AL023807	AL022347	106973 U89959
22550	42224	91414	3 452	7 444	143029			397		36734	41171	32437	157658	1191	157658	6154	999	1 6154	168111	85785	106973
GB_BA1:MTCY22G8	GB BA1:MLCB33	GB_BA1.ECOUW85	GB_EST14.AA448146 452	GB_EST17.AA641937 444	GR PR3.AC003074	GB_BA1:SC1A6	GB_PR4.AC005553	GB_EST3:R49746		GB_BA1:SC6G10	GB_BA1:U00010	GB_BA1:MTCY336	GB_HTG3:AC010579 157658 AC010579	GB_GSS3:B09839	GB_HTG3:AC010579	GB_BA1:SCSECYDN 6154	GB_EST32:AI731596	GB_BA1.SCSECYDN 6154	A GB_PR3:HS525L6	GB_PL2:ATF21P8	GB_PL2:U89959
2358			927			1179) : :			1407			096			1044			1197		
rxa02085			rxa02093			xa02106				rxa02111			xa02112			ra02134			xa02135		

	3-Nov-98	7-Nov-98	26-Jun-98	17-Jun-98	15-Jun-96	15-Jun-96	1-Jul-98	2-Jul-97	g	1-Jul-98	2-Jul-97	25-Jul-96	25-Jul-96	1-Jul-98	15-Jun-96	1-Jul-98
	34,123	31,260	34,281	62,904	36,648	36,648	99,104	99,224	100,000	98,551	98,477	100,000	99,767	99,378	55,504	100,000
	Arabidopsis thaliana	Arabidopsis thaliana	Arabidopsis thaliana	Mycobacterium tuberculosis	Mycobacterium leprae	Mycobacterium leprae	Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Mycobacterium leprae	Corynebacterium glutamicum
Table 4 (continued)	Arabidopsis thaliana chromosome II BAC T3A4 genomic sequence, complete. Arabidopsis thaliana	sequence. Arabidopsis thaliana chromosome 1 BAC F15K9 sequence, complete	sequence. Arabidonsis thaliana BAC T7123, complete sequence.	Mycobacterium tuberculosis H3/Rv complete genome; segment 98/162.	Mycobacterium leprae cosmid B1554 DNA sequence.	Mycobacterium leprae cosmid B1551 DNA sequence.	Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylomithine transaminase (argD), ornithine carbamoyltransferase (argF), arginine repressor (argR), argininosuccinate synthase (argG), and argininosuccinate lyase (argH) genes, complete cds.	Corynebacterium glutamicum N-acetylglutamate-5-semialdehyde dehydrogenase (arqC) gene, complete cds.	C.glutamicum argC, argJ, argB, argD, and argF genes.	Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylornithine transaminase (argD), ornithine carbamoyltransferase (argF), arginine repressor (argR), argininosuccinate synthase (argG), and argininosuccinate lyase (argH) genes, complete cds.	Cognimication of the state of t	C.glutamicum argC, argJ, argB, argD, and argF genes.	C.glutamicum argC, argJ, argB, argD, and argF genes.	Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylornithine transaminase (argD), ornithine carbamoyltransferase (argF), arginine repressor (argR), argininosuccinate synthase (argG), and argininosuccinate lyase (argH) genes, complete cds.	Mycobacterium leprae cosmid B1133 DNA sequence.	Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylornithine transaminase (argD), ornithine carbamoyltransferase (argF), argininosuccinate synthase (argG), and argininosuccinate lyase (argH) genes, complete cds.
	AC005819	AC005278	1189959	270283	L78814	L78813	AF049897	AF005242	X86157	AF049897	AF005242	X86157	X86157	AF049897	L78811	AF049897
	9 57752	71097	106973	34150	36548	36548	9196	1044	4355	9196	1044	4355	4355	9196	C 42106	9196
	GB_PL2:ATAC005819 57752	GB_PL2:F15K9	CB D12-1180059	GB_BA1:MTCY190	GB_BA1:MSGB1554C 36548 S	GB_BA1:MSGB1551C 36548 S	GB_BA2:AF049897	GB_BA1:AF005242	GB_BA1:CGARGCJB 4355 D	GB_BA2:AF049897	GB_BA1:AF005242	GB_BA1:CGARGCJB 4355 D	GB_BA1:CGARGCJB 4355 D	GB_BA2:AF049897	GB_BA1:MSGB1133C 42106 S	GB_BA2:AF049897
	645			1962			903			414			1287			1074
	rxa02136			rxa02139			rxa02153			rxa02154			rxa02155			ra02156

]	121			
25-Jul-96	2-Jun-99 1-Jul-98	25-Jul-96 17-Jun-98	1-Jul-98	5-Jan-99 25-Jul-96	1-Jul-98	5-Jan-99	1-Jul-98	19-Nov-97 22-Apr-96 1-Jul-98
100,000	50.238 99.612	99,612	100,000	99,898	99.843 8 679	100,000	99,774	99,834 rus 65,913 88,524
Corynebacterium	glutamicum Thermotoga maritima Corynebacterium glutamicum	Corynebacterium glutamicum Mycobacterium tuberculosis	Corynebacterium glutamicum	Corynebacterium glutamicum Corynebacterium	glutamicum Gorynebacterium glutamicum	Gorynebacterium glutamicum Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum Streptomyces clavuligerus Corynebacterium glutamicum
Table 4 (continued) C. glutamicum argC, argJ, argB, argD, and argF genes.	Thermotoga maritima section 128 of 136 of the complete genome. Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylornithine transaminase (argD), ornithine carbamoyltransferase (argF), arginine repressor (argR), argininosuccinate synthase (argG), and argininosuccinate lyase (argH) genes, complete cds.	C.glutamicum argC, argJ, argB, argD, and argF genes. Mycobacterium tuberculosis H37Rv complete genome; segment 73/162.	Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylornithine transaminase (argD), ornithine carbamoyltransferase (argF), arginine repressor (argR), argininosuccinate synthase (argG), and arqininosuccinate lyase (argH) genes, complete cds.	Corynebacterium glutamicum ornithine carbamolytransferase (argF) gene, complete cds. Colulamicum ardC, ardJ, ardB, ardD, and ardF genes.	Cigiutamicum argC, argJ, argD, and argP genes. Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylornithine transaminase (argD), ornithine carbamoyltransferase (argF), arginine repressor (argR), argininosuccinate synthase (argG), and argininosuccinate lyase (argH) genes, complete cds.	Corynebacterium giutamicum ornitnine carbamoyiransierase (aigr.) gene, complete cds. Corynebacterium giutamicum arginine repressor (argR) gene, complete cds.	Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylornithine transaminase (argD), ornithine carbamoyltransferase (argF), arginine repressor (argR), argininosuccinate synthase (argG), and argininosuccinate lyase (argH) genes, complete cds.	Corynebacterium glutamicum argininosuccinate synthetase (argG) gene, complete cds. S.davuligerus argG gene and argH gene (partial). Corynebacterium glutamicum N-acetylglutamylphosphate reductase (argC), ornithine acetyltransferase (argJ), N-acetylglutamate kinase (argB), acetylornithine transaminase (argD), ornithine carbamoyltransferase (argF), argininosuccinate synthase (argC), argininosuccinate synthase (argC), and argininosuccinate lyase (argH) genes, complete cds.
X86157	AE001816 AF049897	X86157 Z85982	AF049897	AF031518 X86157	A60197	AF031518 AF041436	AF049897	AF030520 Z49111 AF049897
4355	10007 9196	4355	9196	2045	9196	2045 516	9196	1206 1909 9196
GB_BA1:CGARGCJB 4355	D GB_BA2:AE001816 GB_BA2:AF049897	GB_BA1:CGARGCJB 4355 D GB_BA1:MTCY06H11 38000	GB_BA2:AF049897	GB_BA2.AF031518 2045	GB_BA2.AF049897	GB_BA2:AF031518 GB_BA2:AF041436	GB_BA2.AF049897	GB_BA2.AF030520 GB_BA1.SCARGGH GB_BA2.AF049897
	1296		1080		636		1326	1554
	ma02157		xa02158		жа02159		م302160	ка02162

	1-Jul-98 O.	17-Jun-98	343 86-unr-21	17-Feb-95	19-Jul-97	16-Sep-98	15-Jun-96	16-Sep-98 6-Feb-99	29-Sep-97	1 -Apr-96	22 66-uar-9	6-Feb-99		29-Sep-97	8-Feb-99	•	5-Aug-98		28-Feb-95	28-Feb-95 17-Jun-98	•					28-Feb-95 17-Jun-98 17-Jun-98 17-MAR- 1994 17-Jun-98		
	87,561 1-	64,732 17	36,998 17	39,910 17	38,474 19		40,286 15	33,689 16 99,353 6-	99,367	37 651		93.805		100,000	100,000		39,075		35,542	35,542 33,938	35,542 33,938 65,517	35,542 33,938 65,517 36,770	35,542 33,938 65,517 36,770	35,542 33,938 65,517 36,770 38,674	35,542 33,938 65,517 36,770 38,674 65,465	35,542 33,938 65,517 36,770 38,674 65,465	35,542 33,938 65,517 36,770 38,674 65,465 37,577	35,542 33,938 65,517 36,770 38,674 65,465 37,577 59,823
	Corynebacterium	glutamicum Mycobacterium tuberculosis	Mycobacterium tuberculosis	Corynebacterium	basidiomycete CECT 20197	Homo sapiens	Mycobacterium leprae	Homo sapiens Corynebacterium	glutamicum Corynebacterium	glutamicum	Corynebacterium	glutamicum	alutamicum	Corynebacterium	glutamicum Corynebacterium	glutamicum	Eubacterium	and and an	acidariillopiilluiil Drosophila melanogaster	actidariii lopiii luiti Drosophila melanogaster Mycobacterium tuberculosis	activarimophinam Drosophila melanogaster Mycobacterium tuberculosis Mycobacterium leprae	actoanimophinani Drosophila melanogaster Mycobacterium Mycobacterium leprae Mycobacterium leprae	actoanimoprimari Drosophila melanogaster Mycobacterium leprae Mycobacterium leprae	actorininophinan Drosophila melanogaster Mycobacterium Mycobacterium leprae Mycobacterium leprae	actoantinophinan Drosophila melanogaster Mycobacterium Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae	actualminophinan Drosophila melanogaster Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae	actoantinophinan Drosophila melanogaster Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae	actoantinophinan Drosophila melanogaster Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium leprae Mycobacterium
Table 4 (continued)	Corynebacterium glutamicum argininosuccinate Iyase (argH) gene, complete	cds. Mycobacterium tuberculosis H37Rv complete genome; segment 73/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 41/162.	C.glutamicum glt gene for citrate synthase and ORF.	Basidiomycete CECT 20197 phenoloxidase (pox1) gene, complete cds.	Human Chromosome 15q26.1 PAC clone pDJ417d7, complete sequence.		Human Chromosome 15q26.1 PAC clone pDJ417d7, complete sequence. Brevibacterium flavum aspA gene for aspartase, complete cds.	DNA encoding Brevibacterium flavum asparlase.		Escherichia coli K-12 chromosomal region from 92.8 to 00.1 minutes. Corynebacterium dlutamicum ATP phosphoribosyltransferase (hisG) gene.		Brevibacterium flavum aspA gene for aspariase, comprete cus.	DNA encoding part of aspartase from coryneform bacteria.			Eubacterium acidaminophilum grdR, grdl, grdH genes and partial ldc, grdT		genes. 6t ft. CTS Dm1030 clane DS06059 T7	genes. fruit fly STS Dm1930 clone DS06959 T7. Mycobacterium tuberculosis H37Rv complete genome; segment 95/162.					0, L		0.42	
	AF048764	Z8598Z	Z73101	X66112	U65399	AC002468		AC002468 D25316	E04307		U14003 AF050166		D25316	E08649	AE086704	40 0000 D4	Y17145		2,700	G01195 Z97559	G01195 Z97559	G01195 Z97559 AL035310 U00017	G01195 Z97559 AL035310 U00017	G01195 297559 AL035310 U00017	G01195 Z97559 AL035310 U00017 AL035310	G01195 Z97559 AL035310 U00017 AL035310 Z97559	G01195 Z97559 AL035310 U00017 AL035310 Z97559 U00017	332 G01195 27322 Z97559 40245 AL035310 42157 U00017 42157 U00017 40245 AL035310 27322 Z97559 42157 U00017
	1437		37630	3013	2700	115888	39399	115888	2 8	-	338534	3	1987	188	5 6	407	6019		0	332 27322	332 27322	332 27322 40245 42157	332 27322 40245 42157	332 27322 40245 42157	332 27322 40245 42157 42157	332 27322 40245 42157 42157 27322	332 27322 40245 42157 42157 40245 27322 42157	332 27322 40245 42157 42157 40245 27322 42157
	GB BA2:AF048764	GB_BA1:MTCY06H11 38000	GB_BA1:MTCY31	GB_BA1.CGGLTG	GB_PL2:PGU65399	8970000	GB_BA1:MSGB1970C 39399	S GB_PR3.AC002468 GB_BA1.BB1ASPA	GB_BA1.ENLASI	GB_PAT.E04307	GB_BA1:ECOUW93	GB_BAZ.At CSC SQ	GB_BA1:BRLASPA	0.000 T. F. 40. 000	GB_PA1.E08049	GB_BAZ:AFU86704	GB BA1 EAY17145			GB_STS:G01195 GB_BA1:MTCY261	GB_STS:G01195 GB_BA1:MTCY261	GB_STS:G01195 GB_BA1:MTCY261 GB_BA1:MLCB2533 GB_BA1:U00017	GB_STS:G01195 GB_BA1:MTCY261 GB_BA1:MLCB2533 GB_BA1:U00017	GB_STS:G01195 GB_BA1:MTCY261 GB_BA1:MLCB2533 GB_BA1:U00017 GB_BA1:U00017	GB_STS:G01195 GB_BA1:MLCB2533 GB_BA1:U00017 GB_BA1:U00017	GB_STS:G01195 GB_BA1:MLCB2533 GB_BA1:U00017 GB_BA1:U00017 GB_BA1:MLCB2533 GB_BA1:MLCB2533	GB_STS:G01195 GB_BA1:MTCY261 GB_BA1:U00017 GB_BA1:U00017 GB_BA1:MTCR2533 GB_BA1:MTCR261 GB_BA1:MTCY261	GB_STS:G01195 GB_BA1:MTCY261 GB_BA1:U00017 GB_BA1:U00017 GB_BA1:MTCY261 GB_BA1:MTCY261 GB_BA1:MTCY261
			1251			*30	- 08	101	5		ú	906				393				551	551		551					
			rxa02176			0	rxa02189	2000	rxa02193		0	rxa02194				rxa02195				rxa02197	rxa02197	rxa02197	rxa02197	rxa02197	7802197 7802198	rxa02197	rxa02198	rxa02197

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	29-DEC- 1998	03-DEC-	1996 17-Jun-98		01-MAR- 1994	15-Jun-96	18-Jun-98	22-DEC-	22-MAR- 1997	1-Sep-99	1-Sep-99	23-Jun-98		5-Nov-98	19-0C1- 1998	23-Jun-98	21-Sep-99	0	21-Sep-99	07-OCT-	1997 (Rel. 52, Created)	05-MAR-	1997	31-IMAR- 1999	03-OCT.	1997 (Rel. 52, Created)
	37,191	53,541	40,407		40,541	66,027	71,723	67,101	. 60,870	37,994	37,994	55,844		41,185	38,515	56,282	36,772		36,772	99,515		63,568	000	000'69	52,909	
	Homo sapiens	Mycobacterium	tuberculosis Mycobacterium	tuberculosis	Mycobacterium leprae	Mycobacterium leprae	Mycobacterium tuberculosis	Mycobacterium bovis	Mycobacterium smegmatis 60,870	Homo sapiens	Homo sapiens	Mycobacterium	tuberculosis	Rhodococcus equi	Mus musculus	Mycobacterium	tuberculosis Homo sapiens		Homo sapiens	Corynebacterium	glutamicum	Streptomyces	pristinaespiralis	d Streptomyces spectabilis	Corynebacterium	ammoniagenes
	3 Homo sapiens chromosome 17, clone hCIT.162_E_12, complete sequence.	2 Mycobacterium tuberculosis sequence from clone y154.	Mycobacterium tuberculosis H37Rv complete genome; segment 121/162.		Mycobacterium leprae cosmid B2235.	Mycobacterium leprae cosmid B937 DNA sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 61/162.	Mycobacterium bovis BCG orotidine-5'-monophosphate decarboxylase (uraA)	gene. Mycobacterium smegmatis carbamoyl phosphate synthetase (pyrAB) gene, partial cds and orotidine 5'-monophosphate decarboxylase (pyrF) gene,		_	unordered pieces. Mycobacterium tuberculosis H37Rv complete genome; segment 62/162.			 AU017763 Mouse two-cell stage embryo cDNA Mus musculus cDNA clone J0744A04 3; mRNA sequence. 	Mycobacterium tuberculosis H37Rv complete genome; segment 62/162.	5 Homo sapiens clone NH0549D18, *** SEQUENCING IN PROGRESS ***, 30		 Homo sapiens clone NH0549D18, *** SEQUENCING IN PROGRESS ***, 30 unordered pieces. 	gDNA encoding S-adenosylmethionine synthetase.		Sequence 1 from Patent WO9408014.		4 Streptomyces spectabilis flavoprotein homolog Utp (dtp) gene, partial cds, and Streptomyces spectabilits Stadenosylmethionine synthetase (metK) gene, complete cds.		
	127593 AC006/36	AD0000 02	Z98209		U00019	L78820	Z81011	U01072	U91572	AC009 6	AC009: 6	Z80108		AF077:24	AU017 63	Z80108	AC010 45		AC010 45	E0985		A37831		AF117,74	AB003 93	
	127593	40221	13935		36033	38914	20431	4393	096	192791	192791	39150		5228	286	39150	193862		193862	1239		5392	,	2303	5589	
	GB_PR4:AC006236	GB_BA1:MSGY154	GB BA1:MTCY154	1	GB_BA1:U00019	GB_BA1:MSGB937C	GB_BA1:MTCY2B12	GB_BA2:U01072	GB_BA1:MSU91572	GB_HTG3:AC009364 192791 AC009	GB_HTG3:AC009364 192791 AC009:64	GB_BA1:MTCY21B4	ı	GB_BA2:AF077324 5228	GB_EST22:AU017763	GB_BA1:MTCY21B4 39150	GB_HTG3:AC010745 193862		GB_HTG3:AC010745 193862	EM_PAT.E09855		GB_PAT:A37831		GB_BA2:AF117274	EM_BA1:AB003693	
		948				3462			727			693				1389				1344					1107	
		xa02229				rxa02234			ra02235			xa02237				xa02239				xa02240					rxa02246	

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					Table 4 (continued)		000	70 203 00
		GB_PAT:E07957	5589	E07957	DNA encoding at least guanosine triphosphate cyclonydrolase and riboliavin	Corynebacterium	56,303	16-dap-67
		CB DAT-130742	5589	132742	Symmose: Sequence 1 from patent US 5589355.	Unknown.	52,909	6-Feb-97
77000	334		2689	132743	Sequence 2 from patent US 5589355.	Unknown.	57,937	6-Feb-97
1×907741	00.7	593	5589	AB003693	A for rib operon, complete cds.	Corynebacterium	57,937	03-OCT-
						ammoniagenes		1997 (Rel.
								52, Created)
		GR PAT-132742	5589	132742	Sequence 1 from patent US 5589355.	Unknown	57,937	6-Feb-97
872000	1380	GR PAT-132742	5589	132742	Sequence 1 from patent US 5589355.	Unknown.	61,843	6-Feb-97
04770841	2	EM BA1 ABOO3693	5589	AB003693	Corynebacterium ammoniagenes DNA for rib operon, complete cds.	Conynebacterium	61,843	03-OCT-
						ammoniagenes		1997 (Rel.
							61 043	26, Olcarcu)
		GB_PAT:E07957	5589	E07957	gDNA encoding at least guanosine triphosphate cyclonydrolase and ribollavin. Corylledacterium synthase.	ourynebacierium ammoniagenes	C + 0 - 0	16-dag-67
rxa02249	009-	GB PAT:E07957	5589	E07957	gDNA encoding at least guanosine triphosphate cyclohydrolase and riboflavin	Corynebacterium	64,346	29-Sep-97
		1			synthase.	ammoniagenes	,	
		GB PAT:132742	5589	132742	Sequence 1 from patent US 5589355.	Unknown.	64,346	6-Feb-97
		GB_PAT:132743	2689	132743	Sequence 2 from patent US 5589355.	Unknown.	64,346	6-Feb-97
xa02250	643	GB_PAT_E07957	5589	E07957	gDNA encoding at least guanosine triphosphate cyclohydrolase and riboflavin	Corynebacterium	56,318	29-Sep-97
	!	•			synthase.	ammoniagenes		
		GB PAT-132742	5589	132742	Sequence 1 from patent US 5589355.	Unknown.	56,318	6-Feb-97
		EM_BA1.AB003693	5589	AB003693	Corynebacterium ammoniagenes DNA for rib operon, complete cds.	Corynebacterium	56,318	03-OCT-
						ammoniagenes		1997 (Rel
								52, Created)
CACORY	1269	GB BA1 CGL007732 4460	4460	AJ007732	Corynebacterium glutamicum 3' ppc gene, secG gene, amt gene, ocd gene	Corynebacterium	100,000	7-Jan-99
					and 5' soxA gene.	glutamicum		
		GB BA1;CGAMTGEN 2028	2028	X93513	C.glutamicum amt gene.	Corynebacterium	100,000	29-MAY-
						glutamicum		1996
		GR VIHEHOMVOG	229354	X17403	Human cytomegalovirus strain AD169 complete genome.	human herpesvirus 5	38,651	10-Feb-99
rxa02263	488	GB_BA1:CGL007732	4460		Corynebacterium glutamicum 3' ppc gene, secG gene, amt gene, ocd gene	Corynebacterium	100,000	7-Jan-99
		1			and 5' soxA gene.	glutamicum		
		GB_BA1:CGL007732	4460	AJ007732	Corynebacterium glutamicum 3' ppc gene, secG gene, amt gene, ocd gene and 5' soxA gene.	Corynebacterium glutamicum	37,526	/-Jan-99
rxa02272	1368	EM_PAT:E09373	1591	E09373	Creatinine deiminase gene.	Bacillus sp.	96,928	08-OCT- 1997 (Rel. 52, Created)
		CD BA1-D38505	1591	D38505	Bacillus so nene for creatinine deaminase, complete cds.	Bacillus sp.	96,781	7-Aug-98
rxa02281	1545	GB_GSS12:AQ41101			Homo sapiens, *** SEQUENCING IN PROGRESS ***, 4 unordered pieces. HS_2257_B1_H02_MR CIT Approved Human Genomic Sperm Library D Homo sanians denomic clone Plate=2257 Col=3 Row=P, genomic survey	Homo sapiens Homo sapiens	36,264 36,197	20-Feb-99 17-MAR- 1999
		D.			sequence.			3 :

		GB_EST23;AI128623 363 AI128623 GB_PL2;ATAC007019 102335 AC007019	0 = 4	Homo sapiens Homo sapiens Holete Arabidopsis thaliar	37,017	05-OCT- 1998 16-MAR-
rxa02299	531	GB_BA2:AF116184 540 AF116184 GB GSS9:AQ164310 507 AQ164310	 sequence. 184 Corynebacterium glutamicum L-aspartate-alpha-decarboxylase precursor (panD) gene, complete cds. 310 HS 2171 A2 E01 MR CIT Approved Human Genomic Sperm Library D 	sor Corynebacterium glutamicum D Homo sapiens	100,000	1999 02-MAY- 1999 16-OCT-
ка02311	813	7 878	0 _ 0 0 0		40,288 36,454	3-Sep-96 27-OCT-
		GB_HTG4:AC006091 176878 AC006091	D 4 12	PCI-98 Drosophila meland	36,454	27-OCT- 1999
		GB_BA2:RRU65510 16259 U65510		cool., Rhodospirillum rubrum oxide e activator implete	37,828	9-Apr-97
∝a02315	1752	GB_BA1:MSGY224 40051 AD000004	004 Mycobacterium tuberculosis sequence from clone y224.	Mycobacterium tuberculosis	49,418	03-DEC- 1996
		GB_BA1:MTY25D10 40838 Z95558	Mycobacterium tuberculosis H37Rv complete genome; segment 28/162.		49,360	17-Jun-98
		GB_BA1:MSGY224 40051 AD000004	004 Mycobacterium tuberculosis sequence from clone y224.	Mycobacterium tuberculosis	38,150	03-DEC- 1996
rxa02318	402	GB_HTG3:AC011348 111083 AC011348	348 Homo sapiens chromosome 5 clone CIT-HSPC_303E13, *** SEQUENCING IN PROGRESS ***, 3 ordered pieces.	ICING Homo sapiens	35,821	06-OCT- 1999
		GB_HTG3:AC011348 111083 AC011348		ICING Homo sapiens	35,821	06-OCT- 1999
		GB_HTG3:AC011412 89234 AC011412		ICING Homo sapiens	36,181	06-OCT- 1999
rxa02319	1080	GB_BA1:MSGY224 40051 AD000004	1004 Mycobacterium tuberculosis sequence from clone y224.	Mycobacterium tuberculosis	37,792	03-DEC- 1996
		GB_BA1:MTY25D10 40838 Z95558	Mycobacterium tuberculosis H37Rv complete genome; segment 28/162.	 Mycobacterium tuberculosis 	37,792	17-Jun-98
		GB_EST23:AI117213 476 AI117213	213 ub83h02.r1 Soares 2NbMT Mus musculus cDNA clone IMAGE:1395123 5',mRNA sequence.	23 Mus musculus	35,084	2-Sep-98

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	14-Jan-97	10-Feb-99	10-Feb-99	14-Jan-97		15-Jul-97	1-Nov-95	29-Sep-97	02-DEC- 1994	21-MAY- 1993	2-Aug-96	8-Sep-99	8-Sep-99	17-Jun-98	16-OCT- 1999	16-OCT- 1999	23-Jan-97	17-Jun-98	30 DIA 6	Se-fine-7	9-Sep-94	10-Jun-98	26-Sep-95	10-Jun-99
	61,731	39,624	39,847	64.286	-	36,617	36,617	56,123	56,220	56,220	99,332	36,115	36,115	38,088	35,817	35,817	98,802	38,054	08 520	670,06	100,000	100,000	100,000	39,716
	Corynebacterium	ammoniagenes Mycobacterium	tuberculosis Mycobacterium	tuberculosis Corynebacterium	ammoniagenes	Saccharomyces cerevisiae 36,617	Saccharomyces cerevisiae	unidentified	Unknown.	Unknown.	Corynebacterium glutamicum	Homo sapiens	Homo sapiens	Mycobacterium tuberculosis	Drosophila melanogaster	Drosophila melanogaster	Corynebacterium	glutamicum Mycobacterium	tuberculosis	glutamicum	Corynebacterium glutamicum	Unknown.	Unknown.	Homo sapiens
Table 4 (continued)	B.ammoniagenes purK and purE genes.	Mycobacterium tuberculosis H37Rv complete genome; segment 141/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 141/162.	R ammonianenes purk and purE penes		S.cerevisiae 130kb DNA fragment from chromosome XV.	S.cerevisiae DNA of 51 Kb from chromosome XV right arm.	DNA coding of 2,5-diketogluconic acid reductase.	Sequence 4 from Patent EP 0305608.	Sequence 1 from Patent US 4758514.	Corynebacterium glutamicum Obg protein homolog gene, partial cds, gamma glutamyl kinase (proB) gene, complete cds, and (unkdh) gene, complete cds	Homo sapiens clone NH0012C17, *** SEQUENCING IN PROGRESS ***, 1 unordered pieces.		Mycobacterium tuberculosis H37Rv complete genome, segment 106/162.	Drosophila melanogaster chromosome 3L/75C1 clone RPCI98-3B20, *** SEQUENCING IN PROGRESS ***, 78 unordered pieces.	Drosophila melanogaster chromosome 3L/75C1 clone RPC198-3B20, *** SEQUENCING IN PROGRESS ***, 78 unordered pieces.	C.glutamicum proA gene.	Mycobacterium tuberculosis H37Rv complete genome; segment 107/162.	comments about lightness and sense of the se	Corynebacterium glutamicum Obg protein nomolog gene, partial cus, garrinta glutamyl kinase (proB) gene, complete cds, and (unkdh) gene, complete cds.	C.glutamicum aceA gene and thiX genes (partial).	Sequence 3 from patent US 5700661.	Sequence 3 from patent US 5439822	_ 0,
	X91189	292771	292771	X91189	3	X94335	X90518	E00311	106030	100836	U31230	AC009946	AC009946	Z81368	AC010658	AC010658	X82929	Z81451	0	U31230	X75504	186191	113693	AQ606842
	2582	42729	42729	2582	7007	129528	50984	1853	1853	1853	3005	169072	169072	41230	120754	120754	1783	26914		3005	2427	2135	2135	574
	GB_BA1:BAPURKE	GB_BA1:MTCY71		Ц		GB_PL1:SC130KBXV 129528	GB_PL1:SCXVORFS	GB PAT:E00311	GB_PAT:106030	GB_PAT:100836	GB_BA2:CGU31230	GB_HTG3:AC009946 169072	GB_HTG3:AC009946 169072	GB_BA1:MTCY253	GB_HTG4:AC010658 120754	GB_HTG4:AC010658 120754	GB_BA1:CGPROAGE 1783	N GB BA1:MTCY428		GB_BA2:CGU31230	GB_BA1:CGACEA	GB PAT:186191	GB_PAT:113693	GB_GSS15:AQ60684 2
	1320			010	0			1038	! ! !		1350			777			1419				693			1098
	rxa02345			0300000	1X40 2 330			rxa02373			rxa02375			rxa02380			rxa02382				rxa02400			rxa02432

30-Jun-93 20-Nov-99	7-Feb-99	10-Sep-99	22-Jun-99	17-Jun-98	27-Aug-99	17-Sep-98	00-6nV-7	17-Jun-98	10-MAY-	2-A10-96	oc-fau-z	30-Sep-93	2-Aug-99	17-Jun-98	O1-MAR-	1994	17-Apr-97	28-Jul-98	3-Sep-98	28-Jul-98	17-Jun-98	28-Jul-98	3-Sep-98	04-DEC-	1330 01-MAR.	1994
37,915	100,000	39,175	39,281	39,634	59,343	48,899	C + + + + + + + + + + + + + + + + + + +	59,429	39,510	07 70	641.10	43,249	33,406	39,357	51 76B	5	39,378	39,922	39,922	34,911	54,940	41,265	41,265	37,723	17 77)
Homo sapiens Arabidonsis thallana	Corynebacterium olutamicum		Trypanosoma brucei	Mycobacterium tuberculosis	Mycobacterium leprae	Streptomyces coelicolor	glutamicum	Mycobacterium tuberculosis	Streptomyces coelicolor	Contractor de maria en la contraction de la cont	glutamicum	Neisseria gonorrhoeae	Drosophila melanogaster	Mycobacterium tuberculosis	Mycobacterium lengae	My consected and replace	human herpesvirus 1	Homo sapiens	Homo sapiens	Homo sapiens	Mycobacterium	Homo sapiens	Homo sapiens	Mycobacterium leprae	Mucobactorium langua	אוייים ביים ביים ביים ביים ביים ביים ביים
Table 4 (continued) EST03693 Fetal brain, Stratagene (cat#936206) Homo sapiens cDNA clone HFBDG63 similar to EST containing Alu repeat, mRNA sequence.	Fraction paracteristics and the second of the content of the conte	DDT-0033 Winter flounder ovary Pleuronectes americanus cDNA clone ODT-0033 S' similar to FRUCTOSE-BISPHOSPHATE ALDOLASE B (LIVER), mRNA sequence.	Sheared DNA-5L2.TF Sheared DNA Trypanosoma brucei genomic clone Sheared DNA-5L2, genomic survey sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 84/162.	Mycobacterium leprae cosmid B1788.	Streptomyces coelicolor A3(2) DNA for whiD and whiK loci	Corynebacterium glutamicum (ppx) gene, partial cus.	Mycobacterium tuberculosis H37Rv complete genome; segment 25/162.	Streptomyces coelicolor cosmid E7.	and (Out) and bring a self of the self of	Corynebacterium glutamicum L-proline:INADF+ 3-0xidoreduciase (proc.) gene, corynebacterium complete cd	Neisseria gonorrhoeae pilA gene.	Drosophila melanogaster chromosome 3 clone BACR05C10 (D781) RPCI-98 05.C.10 map 97D-97E strain y; cn bw sp. *** SEQUENCING IN PROGRESS *** 87 unordered pieces.	Mycobacterium tuberculosis H37Rv complete genome; segment 25/162.	Afternation of the Board of the	Mycobacterium lepiae cusmiu bz roo.	Herpes simplex virus (HSV) type 1 complete genome.	Homo sapiens chromosome 19, cosmid R26660, complete sequence.	Homo sapiens chromosome 19, cosmid R26634, complete sequence.	Homo sapiens chromosome 19, cosmid R26660, complete sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 25/162.	Homo sapiens chromosome 19, cosmid R26660, complete sequence.	Homo sapiens chromosome 19, cosmid R26634, complete sequence.	Mycobacterium leprae cosmid L536.		Mycobacterium leprae cosmid B 1490.
T05804	AEU00033	AW013061	AQ650027	Z83859	AL008609	AJ010601	U31224	277162	AL049819		031225	X13965	AC007984	277162	94000	000018	X14112	AC005328	AC005545	AC005328	Z77162	AC005328	AC005545	Z 99125		510000
406	1852	578	728	36021	39228	4692	4.77	37218	16911	,	181/	1920		37218	*000	16674	152261	35414	43514	35414	37218	35414	43514	36224		35881
GB_EST1:T05804	GB_BA2:AF114233	GB_EST37:AW01306	GB_GSS15.AQ65002 728 7	GB_BA1.MTCY359	GB_BA1:MLCB1788	GB_BA1:SCAJ10601	GB_BA2:CGU31224	GB_BA1:MTCY20G9	GB_BA1:SCE7		GB_BA2:CGU31225	GB BAT:NG17PILA	GB_HTG2:AC007984	GB_BA1:MTCY20G9	010000	GB_BA1:000018	GB VI:HE1CG	GB_PR3.AC005328	GB_PR3.AC005545	GB_PR3:AC005328	GB_BA1:MTCY20G9	GR PR3.AC005328	GB_PR3.AC005545	GB_BA1:MLCL536		GB_BA1:000013
	1413			1554		!	1050				933			1188				522			681			1386		
	rxa02458			xa02469			xa02497				rxa02499			rxa02501				rxa02503			rxa02504			rxa02516		

					Table 4 (continued)		,	; ;
		GB_BA1:MTV007	32806	AL021184	Mycobacterium tuberculosis H37Rv complete genome; segment 64/162.	Mycobacterium tuberculosis	61,335	17-Jun-98
rxa02517	920	GB_BA1:MLCL536	36224	299125	Mycobacterium leprae cosmid L536.	Mycobacterium leprae	37,018	04-DEC-
		GB BA1-1100013	35881	100013	Mycobacterium leprae cosmid B1496.	Mycobacterium leprae	37,018	01-MAR-
								1994
		GB BA1:SCC22	22115	AL096839	Streptomyces coelicolor cosmid C22.	Streptomyces coelicolor	37,071	12-Jul-99
rxa02532	1170	GB_OV:AF137219	831	AF137219	ike protein (MII) gene, partial cds.	Amia calva	36,853	7-Sep-99
		GB_EST30:AI645057	301	AI645057	Je	Mus musculus	41,860	29-Apr-99
		GB_EST20:AA822595 429	429	AA822595	37311 Mus musculus cDNA clone	Mus musculus	42,353	17-Feb-98
rxa02536	879	GB_HTG2:AF130866 118874 AF130866	118874	AF130866	Homo sapiens chromosome 8 clone PAC 172N13 map 8q24, *** SEQUENCING IN PROGRESS ***, in unordered pieces.	Homo sapiens	40,754	21-MAR- 1999
		GB_HTG2:AF130866	118874	AF130866	sapiens chromosome 8 clone PAC 172N13 map 8q24, *** ENCING IN PROGRESS *** in unordered pieces	Homo sapiens	40,754	21-MAR- 1999
		GB_PL1:ATT12J5	84499	AL035522	Arabidopsis thaliana DNA chromosome 4, BAC clone T12J5 (ESSAII project). Arabidopsis thaliana	Arabidopsis thaliana	35,063	24-Feb-99
rxa02550	1434	GB_BA1:MTCY279	9150	297991	Mycobacterium tuberculosis H37Rv complete genome; segment 17/162.	Mycobacterium tuberculosis	37,773	17-Jun-98
		GB_BA1:MSGB1970C 39399	39399	L78815	Mycobacterium leprae cosmid B1970 DNA sequence.	Mycobacterium leprae	39,024	15-Jun-96 8
		GB_BA2:SC2H4	25970	AL031514	Streptomyces coelicolor cosmid 2H4.	Streptomyces coelicolor A3(2)	37,906	19-OCT- 1999
rxa02559	1026	GB_BA1:MTV004	69350	AL009198	Mycobacterium tuberculosis H37Rv complete genome; segment 144/162.	Mycobacterium tuberculosis	47,358	18-Jun-98
		GB_PAT:128684 GB_BA1:MTU27357	5100 5100	128684 U27357	Sequence 1 from patent US 5573915. Mycobacterium tuberculosis cyclopropane mycolic acid synthase (cma1)	Unknown. Mycobacterium	39,138 39,138	6-Feb-97 26-Sep-95
		ı			gene, complete cds.	tuberculosis		
rxa02622	1683	GB_BA2:AE001780 GB_OV:AF064564	11997 49254	AE001780 AF064564	Thermotoga maritima section 92 of 136 of the complete genome. Fugu rubripes neurofibromatosis type 1 (NF1), A-kinase anchor protein (AKAP84), BAW protein (BAW), and WSB1 protein (WSB1) genes, complete	Thermotoga maritima Fugu rubripes	44,914 39,732	2-Jun-99 17-Aug-99
		GB_OV:AF064564	49254	AF064564	cds. Fugu rubripes neurofibromatosis type 1 (NF1), A-kinase anchor protein (AKAP84), BAW protein (BAW), and WSB1 protein (WSB1) genes, complete	Fugu rubripes	36,703	17-Aug-99
∝a02623	714	GB_GSS5:AQ818728 444	444	AQ818728	USS. 5268_A1_G09_SP6E RPCI-11 Human Male BAC Library Homo sapiens genomic clone Plate=844 Col=17 Row=M, genomic survey sequence.	Homo sapiens	38,801	26-Aug-99
		GB_HTG5:AC011083 198586	198586	AC011083		Homo sapiens	35,714	19-Nov-99
		GB_GSS6:AQ826948 544	544	AQ826948	HS_5014_A2_C12_T7A RPCI-11 Human Male BAC Library Homo sapiens genomic clone Plate=590 Col=24 Row=E, genomic survey sequence.	Homo sapiens	39,146	27-Aug-99

																_		
28-Apr-93	28-Apr-93	07-MAR- 1997	07-MAR-	26-Apr-93	26-Apr-93	07-MAR- 1997	07-MAR- 1997	24-MAY- 1993	20-MAR- 1997	27-Jun-98		26-Apr-93	29-Sep-97	29-Sep-97	25-MAR- 1999	22-Nov-99	28-MAY- 1998	2-Aug-99
137,013	137,013	39,130	39,130	39,130	99,138	990'66	990'66	38,402	38,655	36,074		99,715	98,523	98,523	36,593	36,089	36,089	32,757
Bovine respiratory syncytial 37,013 virus	Bovine respiratory syncytial 37,013 virus	Corynebacterium	graamicum Corynebacterium alutamicum	grutamicum Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Ictalurus punctatus	Mus musculus	Homo sapiens		Corynebacterium glutamicum	Corynebacterium glutamicum	Corynebacterium glutamicum	Hordeum vulgare	Homo sapiens	Homo sapiens	Drosophila melanogaster
Table 4 (continued) Sovine respiratory syncytial virus membrane glycoprotein mRNA, complete	Sovine respiratory syncytial virus membrane glycoprotein mRNA, complete ds.	Sequence 1 from Patent WO9519442.	Sequence 5 from Patent WO9519442.	Sorynebacterium glutamicum threonine dehydratase (ilvA) gene, complete des.	Dorynebacterium glutamicum threonine dehydratase (ilvA) gene, complete ;ds.	Sequence 9 from Patent WO9519442.	Sequence 7 from Patent WO9519442.	ctalurus punctatus cyclic nucleotide-gated channel RNA sequence.	mx91c06.r1 Soares mouse NML Mus musculus cDNA clone IMAGE.6937065. mRNA sequence.	CIT-HSP-2294E14.TR CIT-HSP Homo sapiens genomic clone 2294E14, penomic survey sequence.		C.glutamicum pheA gene encoding prephenate dehydratase, complete cds.	DNA encoding prephenate dehydratase.	DNA encoding prephenate dehydratase.	Hordeum vulgare DNA for chromosome 4H	Human DNA sequence from cosmid 310H5 from a contig from the tip of the short arm of chromosome 16, spanning 2Mb of 16p13.3. Contains EST and CnG island	Homo sapiens chromosome 16, cosmid clone RT286 (LANL), complete sequence.	Drosophila melanogaster chromosome 3 clone BACR16118 (D815) RPCI-98 16.1.18 map 95A-95A strain y; cn bw sp, *** SEQUENCING IN PROGRESS***, 101 unordered pieces.
M86652	M86652	A45577	A45581	L01508	L01508	A45585	A45583	M83111	AA265464	AQ006950		M13774	E04483	E06110	Y14573	269705	AC004754	AC008223
462	462	1925	1925	1925	1925	1925	1925	2049	4 345	480		1088	948	948	59748	29718	39188	3 130212
GB_VI:BRSMGP	GB_VI:BRSMGP	GB_PAT:A45577	GB_PAT:A45581	GB_BA1.CORILVA	GB_BA1:CORILVA	GB_PAT:A45585	GB_PAT:A45583	GB_OV:ICTCNC	GB_EST11:AA265464 345	GB_GSS8:AQ006950 480		GB_BA1:CORPHEA	GB_PAT:E04483	GB_PAT:E06110	GB_PL1:HVCH4H	GB_PR2:HS310H5	GB_PR3.AC004754	GB_HTG2.AC008223 130212 AC008223
708		1953			1392			1326				1068			1005			1461
rxa02629		rxa02645			rxa02646			xa02648			∝a02653	rxa02687			xa02717			rxa02754

	2-Aug-99	10-Feb-99	9-Nov-99	5-Nov-99	20-Jan-99	14-Sep-98	04-DEC- 1996	22-Jul-99	14-Sep-98	17-Jun-98	01-MAR- 1994	17-Jun-98	3-Jun-99	3-Jun-99	17-Jun-98	28-Apr-93 9-Aug-94	27-OCT- 1998
	32,757	37,838	35,331	33,807	36,929	99,852	43,836	48,588	99,914	38,339	38,996	37,640	37,906	35,280	39,765	38,937 s 38,495	40.828
	Drosophila melanogaster	Mycobacterium tuberculosis	Homo sapiens	Homo sapiens	Burkholderia pseudomallei 36,929	Corynebacterium glutamicum	Caenorhabditis elegans	Caenorhabditis elegans	Corynebacterium glutamicum	Mycobacterium tuberculosis	Mycobacterium leprae	Mycobacterium tuberculosis	Homo sapiens	Homo sapiens	Mycobacterium tuberculosis	Gallus gallus Mycobacterium smegmatis	Homo sapiens
Table 4 (continued)	Drosophila melanogaster chromosome 3 clone BACR16118 (D815) RPCI-98 16.1.18 map 95A-95A strain y; cn bw sp. *** SEQUENCING IN PROGRESS ***, 101 unordered pieces.	Mycobacterium tuberculosis H37Rv complete genome; segment 141/162	Homo sapiens clone 14_B_7, *** SEQUENCING IN PROGRESS ***, 20 unordered pieces.	Homo sapiens clone 14_B_7, *** SEQUENCING IN PROGRESS ***, 20 unordered pieces.	Burkholderia pseudomallei putative dihydroorotase (pyrC) gene, partial cds, putative 1-acyl-sn-glycerol-3-phosphate acyltransferase (plsC), putative diadenosine tetraphosphatase (apaH), complete cds; type II O-antigen biosynthesis gene cluster, complete sequence; putative undecaprenyl phosphate N-acetylglucosaminyltransferase, and putative UDP-glucose 4-epimerase genes, complete cds; and putative galactosyl transferase gene,	partial cos. Corynebacterium glutamicum dipeptide-binding protein (dciAE) gene, partial cos; adenine phosphoribosyltransferase (apt) and GTP pyrophosphokinase cos; complete core; and unknown nene	Caenorhabditis elegans cosmid T19B4.	AV193572 Yuji Kohara unpublished cDNA;Strain N2 hermaphrodite embryo Caenorhabditis elegans cDNA clone yk618h8 5', mRNA sequence.	Corynebacterium glutamicum dipeptide-binding protein (dciAE) gene, partial cds; adenine phosphoribosyltransferase (apt) and GTP pyrophosphokinase (rel) enes, complete cds, and unknown gene.	Mycobacterium tuberculosis H37Rv complete genome; segment 114/162.	Mycobacterium leprae cosmid B1177.	Mycobacterium tuberculosis H37Rv complete genome; segment 111/162.	Homo sapiens 12p21 BAC RPCI11-259018 (Roswell Park Cancer Institute Human RAC Library) complete seguence	Human BAC Library) complete sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 111/162.	Chicken tyrosine kınase (cek2) mRNA, complete cds. M.smegmatis asd, ask-alpha, and ask-beta genes.	qg48g01.x1 Soares_testis_NHT Homo sapiens cDNA clone IMAGE:1838448 Homo sapiens 3' similar to WP:C25D7.8 CE08394 ;, mRNA sequence.
	AC008223	42729 Z92771	AC011678	AC011678	AF064070	AF038651	U80438	AV193572	AF038651	277724	U00011	Z83863	AC006581	172931 AC006581	Z83863	M35195 Z17372	AI223401
	130212	42729	171967	171967	23183	4077	37121	360	4077	35946	40429	33818	172931	172931	33818	3694 5037	169
	GB_HTG2:AC008223 130212 AC008223	GB_BA1:MTCY71	GB_HTG5:AC011678 171967	GB_HTG5:AC011678	GB_BA2:AF064070	GB_BA2:AF038651	GB_IN1:CELT19B4	GB_EST36:AV193572 360	GB_BA2:AF038651	GB_BA1:MTCY227	GB_BA1:U00011	GB_BA1:MTCY159	GB_PR4:AC006581	GB_PR4:AC006581	GB_BA1:MTCY159	GB_OV:CHKCEK2 GB_BA1:MSASDASK	GB_EST24;Al223401 169
			1422			678			1158			1266			951		1194
			rxa02758			xa02771			rxa02772			rxa02733			rxa02791		xa02802

27-OCT-	1998	17-Jun-98	17-Jun-98	8-Jan-98	17-Jun-98	17-Jun-98	09-MAR- 1995	25-Apr-96	27-MAY-	1998	30-Sep-98	15-DEC- 1999	17-DEC-	17-DEC- 1999
40 828		58,418	40,496	39,826	100.000	37,710	39,626	icur88,854	41,489	-	38,005	39,869	34,930	34,634
Homo saniens		Mycobacterium tuberculosis	Mycobacterium tuberculosis	Homo sapiens	Corynebacterium glutamicum	Mycobacterium tuberculosis	Mycobacterium leprae	Corynebacterium glutamicum 8,854	Mus musculus		Mus musculus	Leishmania major	Homo sapiens	Homo sapiens
Table 4 (continued)	gg4sg01.x1.soares_tests_nn1 norto saprens conscioned in social process similar to WP.C25D7.8 CE08394 ;; mRNA sequence.	Mycobacterium tuberculosis H37Rv complete genome; segment 138/162.	Mycobacterium tuberculosis H37Rv complete genome; segment 138/162.	Homo sapiens mRNA for hB-FABP.	Corynebacterium glutamicum dapD gene, complete CDS.	Mycobacterium tuberculosis H37Rv complete genome; segment 52/162.	Aycobacterium leprae cosmid B1756.	Blactofermentum orf1 gene and sig8 gene.	1932912 rt Soares mammary pland NhMMG Mus musculus cDNA clone	MAGE:1348414 5' similar to TR:Q61025 Q61025 HYPOTHETICAL 15.2 KD PROTFIN	d27c05.r1 Soares_thymus_2NbMT Mus musculus cDNA clone MAGE:1447112 5', mRNA sequence.	eishmania major Friedlin chromosome 4 cosmid L2743.	duman DNA sequence from clone RP1-61B2 on chromosome 6p11.2-12.3 Contains isoforms 1 and 3 of BPAG1 (bullous pemphigoid antigen 1 230/240kD), an exon of a gene similar to murine MACF cytoskeletal protein, ETS and CSSs. complete sequence.	First and Coos, Comprete sequence. Human DNA sequence from clone RP1-61B2 on chromosome 6p11.2-12.3 Contains isoforms 1 and 3 of BPAG1 (bullows pemphigoid antigen 1 230/240kD), an exon of a gene similar to more MACF cytoskeletal protein, STSs and GSSs, complete sequence.
10000000	AI223401	295120	Z95120	AJ002962	AJ004934	293777	U15180	749824	A 4 9 8 0 2 3 7	V20062	AI158316	AL031910	AL096710	119666 AL096710
Ç	691	22070	22070	778	1160	29540	38675	2906	277	ò	371	38368	119666	119666
	GB_ES124:Al223401 169	GB_BA1:MTCY7D11	GB_BA1:MTCY7D11	GB_PR1:HSAJ2962	GB_BA1:CGAJ4934	GB_BA1:MTCI364	GB_BA1:MLU15180	CB BA1 BI SICBCN	CD EST13: A A080237 377	GB_ES121.A4360251	GB_EST23:AI158316_371	GB_IN1:LMFL2743	GB_PR3:HSDJ61B2	GB_PR3 HSDJ61B2
		494			809			653	206			1237		
		rxa02814			xa02843			30000	C07C0SX)			xs03223		

Exemplification

Example 1: Preparation of total genomic DNA of Corynebacterium glutamicum ATCC 13032

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A culture of Corynebacterium glutamicum (ATCC 13032) was grown overnight 5 at 30°C with vigorous shaking in BHI medium (Difco). The cells were harvested by centrifugation, the supernatant was discarded and the cells were resuspended in 5 ml buffer-I (5% of the original volume of the culture — all indicated volumes have been calculated for 100 ml of culture volume). Composition of buffer-I: 140.34 g/l sucrose, 10 $2.46 \text{ g/l MgSO}_4 \times 7H_2O$, 10 ml/l KH₂PO₄ solution (100 g/l, adjusted to pH 6.7 with KOH), 50 ml/l M12 concentrate (10 g/l (NH₄)₂SO₄, 1 g/l NaCl, 2 g/l MgSO₄ x 7H₂O, 0.2 g/l CaCl₂, 0.5 g/l yeast extract (Difco), 10 ml/l trace-elements-mix (200 mg/l FeSO₄ x H₂O, 10 mg/l ZnSO₄ x 7 H₂O, 3 mg/l MnCl₂ x 4 H₂O, 30 mg/l H₃BO₃ 20 mg/l CoCl₂ x 6 H₂O, 1 mg/l NiCl₂ x 6 H₂O, 3 mg/l Na₂MoO₄ x 2 H₂O, 500 mg/l complexing agent (EDTA or critic acid), 100 ml/l vitamins-mix (0.2 mg/l biotin, 0.2 mg/l folic acid, 20 mg/l p-amino benzoic acid, 20 mg/l riboflavin, 40 mg/l ca-panthothenate, 140 mg/l nicotinic acid, 40 mg/l pyridoxole hydrochloride, 200 mg/l myo-inositol). Lysozyme was added to the suspension to a final concentration of 2.5 mg/ml. After an approximately 4 h incubation at 37°C, the cell wall was degraded and the resulting protoplasts are harvested by centrifugation. The pellet was washed once with 5 ml 20 buffer-I and once with 5 ml TE-buffer (10 mM Tris-HCl, 1 mM EDTA, pH 8). The pellet was resuspended in 4 ml TE-buffer and 0.5 ml SDS solution (10%) and 0.5 ml NaCl solution (5 M) are added. After adding of proteinase K to a final concentration of 200 μg/ml, the suspension is incubated for ca.18 h at 37°C. The DNA was purified by extraction with phenol, phenol-chloroform-isoamylalcohol and chloroformisoamylalcohol using standard procedures. Then, the DNA was precipitated by adding 1/50 volume of 3 M sodium acetate and 2 volumes of ethanol, followed by a 30 min incubation at -20°C and a 30 min centrifugation at 12,000 rpm in a high speed centrifuge using a SS34 rotor (Sorvall). The DNA was dissolved in 1 ml TE-buffer containing 20 $\mu g/ml$ RNaseA and dialysed at 4°C against 1000 ml TE-buffer for at least 3 hours. During this time, the buffer was exchanged 3 times. To aliquots of 0.4 ml of the dialysed DNA solution, 0.4 ml of 2 M LiCl and 0.8 ml of ethanol are added. After a 30

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min incubation at -20°C, the DNA was collected by centrifugation (13,000 rpm, Biofuge Fresco, Heraeus, Hanau, Germany). The DNA pellet was dissolved in TE-buffer. DNA prepared by this procedure could be used for all purposes, including southern blotting or construction of genomic libraries.

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Example 2: Construction of genomic libraries in *Escherichia coli* of *Corynebacterium glutamicum* ATCC13032.

Using DNA prepared as described in Example 1, cosmid and plasmid libraries were constructed according to known and well established methods (*see e.g.*, Sambrook, J. *et al.* (1989) "Molecular Cloning: A Laboratory Manual", Cold Spring Harbor Laboratory Press, or Ausubel, F.M. *et al.* (1994) "Current Protocols in Molecular Biology", John Wiley & Sons.)

Any plasmid or cosmid could be used. Of particular use were the plasmids pBR322 (Sutcliffe, J.G. (1979) *Proc. Natl. Acad. Sci. USA*, 75:3737-3741); pACYC177 (Change & Cohen (1978) *J. Bacteriol* 134:1141-1156), plasmids of the pBS series (pBSSK+, pBSSK- and others; Stratagene, LaJolla, USA), or cosmids as SuperCos1 (Stratagene, LaJolla, USA) or Lorist6 (Gibson, T.J., Rosenthal A. and Waterson, R.H. (1987) *Gene* 53:283-286. Gene libraries specifically for use in *C. glutamicum* may be constructed using plasmid pSL109 (Lee, H.-S. and A. J. Sinskey (1994) *J. Microbiol. Biotechnol.* 4: 256-263).

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Example 3: DNA Sequencing and Computational Functional Analysis

Genomic libraries as described in Example 2 were used for DNA sequencing according to standard methods, in particular by the chain termination method using ABI377 sequencing machines (see *e.g.*, Fleischman, R.D. *et al.* (1995) "Whole-genome Random Sequencing and Assembly of Haemophilus Influenzae Rd., *Science*, 269:496-512). Sequencing primers with the following nucleotide sequences were used: 5'-GGAAACAGTATGACCATG-3' or 5'-GTAAAACGACGGCCAGT-3'.

Example 4: In vivo Mutagenesis

In vivo mutagenesis of Corynebacterium glutamicum can be performed by passage of plasmid (or other vector) DNA through E. coli or other microorganisms (e.g. Bacillus spp. or yeasts such as Saccharomyces cerevisiae) which are impaired in their capabilities to maintain

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the integrity of their genetic information. Typical mutator strains have mutations in the genes for the DNA repair system (e.g., mutHLS, mutD, mutT, etc.; for reference, see Rupp, W.D. (1996) DNA repair mechanisms, in: *Escherichia col*i and *Salmonella*, p. 2277-2294, ASM: Washington.) Such strains are well known to those of ordinary skill in the art. The use of such strains is illustrated, for example, in Greener, A. and Callahan, M. (1994) <u>Strategies</u> 7: 32-34.

Example 5: DNA Transfer Between Escherichia coli and Corynebacterium glutamicum

Several Corynebacterium and Brevibacterium species contain endogenous plasmids (as e.g., pHM1519 or pBL1) which replicate autonomously (for review see, e.g., 10 Martin, J.F. et al. (1987) Biotechnology, 5:137-146). Shuttle vectors for Escherichia coli and Corynebacterium glutamicum can be readily constructed by using standard vectors for E. coli (Sambrook, J. et al. (1989), "Molecular Cloning: A Laboratory Manual", Cold Spring Harbor Laboratory Press or Ausubel, F.M. et al. (1994) "Current Protocols in Molecular Biology", John Wiley & Sons) to which a origin or replication for and a 15 suitable marker from Corynebacterium glutamicum is added. Such origins of replication are preferably taken from endogenous plasmids isolated from Corynebacterium and Brevibacterium species. Of particular use as transformation markers for these species are genes for kanamycin resistance (such as those derived from the Tn5 or Tn903 transposons) or chloramphenicol (Winnacker, E.L. (1987) "From Genes to Clones — Introduction to Gene Technology, VCH, Weinheim). There are numerous examples in the literature of the construction of a wide variety of shuttle vectors which replicate in both E. coli and C. glutamicum, and which can be used for several purposes, including gene overexpression (for reference, see e.g., Yoshihama, M. et al. (1985) J. Bacteriol. 162:591-597, Martin J.F. et al. (1987) Biotechnology, 5:137-146 and Eikmanns, B.J. et al. (1991) Gene, 25 102:93-98).

Using standard methods, it is possible to clone a gene of interest into one of the shuttle vectors described above and to introduce such a hybrid vectors into strains of *Corynebacterium glutamicum*. Transformation of *C. glutamicum* can be achieved by protoplast transformation (Kastsumata, R. *et al.* (1984) *J. Bacteriol.* 159306-311), electroporation (Liebl, E. *et al.* (1989) *FEMS Microbiol. Letters*, 53:399-303) and in cases where special vectors are used, also by conjugation (as described *e.g.* in Schafer, A *et al.*

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(1990) *J. Bacteriol*. 172:1663-1666). It is also possible to transfer the shuttle vectors for *C. glutamicum* to *E. coli* by preparing plasmid DNA from *C. glutamicum* (using standard methods well-known in the art) and transforming it into *E. coli*. This transformation step can be performed using standard methods, but it is advantageous to use an Mcr-deficient *E. coli* strain, such as NM522 (Gough & Murray (1983) *J. Mol. Biol.* 166:1-19).

Genes may be overexpressed in *C. glutamicum* strains using plasmids which comprise pCG1 (U.S. Patent No. 4,617,267) or fragments thereof, and optionally the gene for kanamycin resistance from TN903 (Grindley, N.D. and Joyce, C.M. (1980) *Proc. Natl. Acad. Sci. USA* 77(12): 7176-7180). In addition, genes may be overexpressed in *C. glutamicum* strains using plasmid pSL109 (Lce, H.-S. and A. J. Sinskey (1994) *J. Microbiol. Biotechnol.* 4: 256-263).

Aside from the use of replicative plasmids, gene overexpression can also be achieved by integration into the genome. Genomic integration in *C. glutamicum* or other Corynebacterium or Brevibacterium species may be accomplished by well-known methods, such as homologous recombination with genomic region(s), restriction endonuclease mediated integration (REMI) (see, *e.g.*, DE Patent 19823834), or through the use of transposons. It is also possible to modulate the activity of a gene of interest by modifying the regulatory regions (*e.g.*, a promoter, a repressor, and/or an enhancer) by sequence modification, insertion, or deletion using site-directed methods (such as homologous recombination) or methods based on random events (such as transposon mutagenesis or REMI). Nucleic acid sequences which function as transcriptional terminators may also be inserted 3' to the coding region of one or more genes of the invention, such terminators are well-known in the art and are described, for example, in Winnacker, E.L. (1987) From Genes to Clones – Introduction to Gene Technology. VCH: Weinheim.

Example 6: Assessment of the Expression of the Mutant Protein

Observations of the activity of a mutated protein in a transformed host cell rely on the fact that the mutant protein is expressed in a similar fashion and in a similar quantity to that of the wild-type protein. A useful method to ascertain the level of transcription of the mutant gene (an indicator of the amount of mRNA available for translation to the gene product) is to perform a Northern blot (for reference see, for example, Ausubel *et al.*

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(1988) Current Protocols in Molecular Biology, Wiley: New York), in which a primer designed to bind to the gene of interest is labeled with a detectable tag (usually radioactive or chemiluminescent), such that when the total RNA of a culture of the organism is extracted, run on gel, transferred to a stable matrix and incubated with this probe, the binding and quantity of binding of the probe indicates the presence and also the quantity of mRNA for this gene. This information is evidence of the degree of transcription of the mutant gene. Total cellular RNA can be prepared from *Corynebacterium glutamicum* by several methods, all well-known in the art, such as that described in Bormann, E.R. *et al.* (1992) *Mol. Microbiol.* 6: 317-326.

To assess the presence or relative quantity of protein translated from this mRNA, standard techniques, such as a Western blot, may be employed (see, for example, Ausubel et al. (1988) Current Protocols in Molecular Biology, Wiley: New York). In this process, total cellular proteins are extracted, separated by gel electrophoresis, transferred to a matrix such as nitrocellulose, and incubated with a probe, such as an antibody, which specifically binds to the desired protein. This probe is generally tagged with a chemiluminescent or colorimetric label which may be readily detected. The presence and quantity of label observed indicates the presence and quantity of the desired mutant protein present in the cell.

20 Example 7: Growth of Genetically Modified Corynebacterium glutamicum — Media and Culture Conditions

Genetically modified *Corynebacteria* are cultured in synthetic or natural growth media. A number of different growth media for Corynebacteria are both well-known and readily available (Lieb *et al.* (1989) *Appl. Microbiol. Biotechnol.*, 32:205-210; von der Osten *et al.* (1998) Biotechnology Letters, 11:11-16; Patent DE 4,120,867; Liebl (1992) "The Genus *Corynebacterium*, in: The Procaryotes, Volume II, Balows, A. *et al.*, eds. Springer-Verlag). These media consist of one or more carbon sources, nitrogen sources, inorganic salts, vitamins and trace elements. Preferred carbon sources are sugars, such as mono-, di-, or polysaccharides. For example, glucose, fructose, mannose, galactose, ribose, sorbose, ribulose, lactose, maltose, sucrose, raffinose, starch or cellulose serve as very good carbon sources. It is also possible to supply sugar to the media via complex compounds such as molasses or other by-products from sugar refinement. It can also be

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advantageous to supply mixtures of different carbon sources. Other possible carbon sources are alcohols and organic acids, such as methanol, ethanol, acetic acid or lactic acid. Nitrogen sources are usually organic or inorganic nitrogen compounds, or materials which contain these compounds. Exemplary nitrogen sources include ammonia gas or ammonia salts, such as NH₄Cl or (NH₄)₂SO₄, NH₄OH, nitrates, urea, amino acids or complex nitrogen sources like corn steep liquor, soy bean flour, soy bean protein, yeast extract, meat extract and others.

Inorganic salt compounds which may be included in the media include the chloride-, phosphorous- or sulfate- salts of calcium, magnesium, sodium, cobalt, molybdenum, potassium, manganese, zinc, copper and iron. Chelating compounds can be added to the medium to keep the metal ions in solution. Particularly useful chelating compounds include dihydroxyphenols, like catechol or protocatechuate, or organic acids, such as citric acid. It is typical for the media to also contain other growth factors, such as vitamins or growth promoters, examples of which include biotin, riboflavin, thiamin, folic acid, nicotinic acid, pantothenate and pyridoxin. Growth factors and salts frequently originate from complex media components such as yeast extract, molasses, corn steep liquor and others. The exact composition of the media compounds depends strongly on the immediate experiment and is individually decided for each specific case. Information about media optimization is available in the textbook "Applied Microbiol. Physiology, A Practical Approach (eds. P.M. Rhodes, P.F. Stanbury, IRL Press (1997) pp. 53-73, ISBN 0 19 963577 3). It is also possible to select growth media from commercial suppliers, like standard 1 (Merck) or BHI (grain heart infusion, DIFCO) or others.

All medium components are sterilized, either by heat (20 minutes at 1.5 bar and 121°C) or by sterile filtration. The components can either be sterilized together or, if necessary, separately. All media components can be present at the beginning of growth, or they can optionally be added continuously or batchwise.

Culture conditions are defined separately for each experiment. The temperature should be in a range between 15°C and 45°C. The temperature can be kept constant or can be altered during the experiment. The pH of the medium should be in the range of 5 to 8.5, preferably around 7.0, and can be maintained by the addition of buffers to the media. An exemplary buffer for this purpose is a potassium phosphate buffer. Synthetic buffers such as MOPS, HEPES, ACES and others can alternatively or simultaneously be used. It

is also possible to maintain a constant culture pH through the addition of NaOH or NH₄OH during growth. If complex medium components such as yeast extract are utilized, the necessity for additional buffers may be reduced, due to the fact that many complex compounds have high buffer capacities. If a fermentor is utilized for culturing the microorganisms, the pH can also be controlled using gaseous ammonia.

The incubation time is usually in a range from several hours to several days. This time is selected in order to permit the maximal amount of product to accumulate in the broth. The disclosed growth experiments can be carried out in a variety of vessels, such as microtiter plates, glass tubes, glass flasks or glass or metal fermentors of different sizes.

For screening a large number of clones, the microorganisms should be cultured in microtiter plates, glass tubes or shake flasks, either with or without baffles. Preferably 100 ml shake flasks are used, filled with 10% (by volume) of the required growth medium. The flasks should be shaken on a rotary shaker (amplitude 25 mm) using a speed-range of 100 – 300 rpm. Evaporation losses can be diminished by the maintenance of a humid atmosphere; alternatively, a mathematical correction for evaporation losses should be performed.

If genetically modified clones are tested, an unmodified control clone or a control clone containing the basic plasmid without any insert should also be tested. The medium is inoculated to an OD₆₀₀ of O.5 – 1.5 using cells grown on agar plates, such as CM plates (10 g/l glucose, 2,5 g/l NaCl, 2 g/l urea, 10 g/l polypeptone, 5 g/l yeast extract, 5 g/l meat extract, 22 g/l NaCl, 2 g/l urea, 10 g/l polypeptone, 5 g/l yeast extract, 5 g/l meat extract, 22 g/l agar, pH 6.8 with 2M NaOH) that had been incubated at 30°C. Inoculation of the media is accomplished by either introduction of a saline suspension of *C. glutamicum* cells from CM plates or addition of a liquid preculture of this bacterium.

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Example 8 - In vitro Analysis of the Function of Mutant Proteins

The determination of activities and kinetic parameters of enzymes is well established in the art. Experiments to determine the activity of any given altered enzyme must be tailored to the specific activity of the wild-type enzyme, which is well within the ability of one of ordinary skill in the art. Overviews about enzymes in general, as well as specific details concerning structure, kinetics, principles, methods, applications and examples for the determination of many enzyme activities may be

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found, for example, in the following references: Dixon, M., and Webb, E.C., (1979)
Enzymes. Longmans: London; Fersht, (1985) Enzyme Structure and Mechanism.
Freeman: New York; Walsh, (1979) Enzymatic Reaction Mechanisms. Freeman: San Francisco; Price, N.C., Stevens, L. (1982) Fundamentals of Enzymology. Oxford Univ.
Press: Oxford; Boyer, P.D., ed. (1983) The Enzymes, 3rd ed. Academic Press: New York; Bisswanger, H., (1994) Enzymkinetik, 2nd ed. VCH: Weinheim (ISBN 3527300325); Bergmeyer, H.U., Bergmeyer, J., Graßl, M., eds. (1983-1986) Methods of Enzymatic Analysis, 3rd ed., vol. I-XII, Verlag Chemie: Weinheim; and Ullmann's Encyclopedia of Industrial Chemistry (1987) vol. A9, "Enzymes". VCH: Weinheim, p.

The activity of proteins which bind to DNA can be measured by several well-established methods, such as DNA band-shift assays (also called gel retardation assays). The effect of such proteins on the expression of other molecules can be measured using reporter gene assays (such as that described in Kolmar, H. *et al.* (1995) *EMBO J.* 14: 3895-3904 and references cited therein). Reporter gene test systems are well known and established for applications in both pro- and eukaryotic cells, using enzymes such as beta-galactosidase, green fluorescent protein, and several others.

The determination of activity of membrane-transport proteins can be performed according to techniques such as those described in Gennis, R.B. (1989) "Pores, Channels and Transporters", in Biomembranes, Molecular Structure and Function, Springer: Heidelberg, p. 85-137; 199-234; and 270-322.

Example 9: Analysis of Impact of Mutant Protein on the Production of the Desired Product

The effect of the genetic modification in *C. glutamicum* on production of a desired compound (such as an amino acid) can be assessed by growing the modified microorganism under suitable conditions (such as those described above) and analyzing the medium and/or the cellular component for increased production of the desired product (*i.e.*, an amino acid). Such analysis techniques are well known to one of ordinary skill in the art, and include spectroscopy, thin layer chromatography, staining methods of various kinds, enzymatic and microbiological methods, and analytical chromatography such as high performance liquid chromatography (see, for example,

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Ullman, Encyclopedia of Industrial Chemistry, vol. A2, p. 89-90 and p. 443-613, VCH: Weinheim (1985); Fallon, A. et al., (1987) "Applications of HPLC in Biochemistry" in: Laboratory Techniques in Biochemistry and Molecular Biology, vol. 17; Rehm et al. (1993) Biotechnology, vol. 3, Chapter III: "Product recovery and purification", page 469-714, VCH: Weinheim; Belter, P.A. et al. (1988) Bioseparations: downstream processing for biotechnology, John Wiley and Sons; Kennedy, J.F. and Cabral, J.M.S. (1992) Recovery processes for biological materials, John Wiley and Sons; Shaeiwitz, J.A. and Henry, J.D. (1988) Biochemical separations, in: Ulmann's Encyclopedia of Industrial Chemistry, vol. B3, Chapter 11, page 1-27, VCH: Weinheim; and Dechow, F.J. (1989) Separation and purification techniques in biotechnology, Noyes Publications.)

In addition to the measurement of the final product of fermentation, it is also possible to analyze other components of the metabolic pathways utilized for the production of the desired compound, such as intermediates and side-products, to determine the overall efficiency of production of the compound. Analysis methods include measurements of nutrient levels in the medium (e.g., sugars, hydrocarbons, nitrogen sources, phosphate, and other ions), measurements of biomass composition and growth, analysis of the production of common metabolites of biosynthetic pathways, and measurement of gasses produced during fermentation. Standard methods for these measurements are outlined in Applied Microbial Physiology, A Practical Approach, P.M. Rhodes and P.F. Stanbury, eds., IRL Press, p. 103-129; 131-163; and 165-192 (ISBN: 0199635773) and references cited therein.

Example 10: Purification of the Desired Product from C. glutamicum Culture

25 Recovery of the desired product from the *C. glutamicum* cells or supernatant of the above-described culture can be performed by various methods well known in the art. If the desired product is not secreted from the cells, the cells can be harvested from the culture by low-speed centrifugation, the cells can be lysed by standard techniques, such as mechanical force or sonication. The cellular debris is removed by centrifugation, and the supernatant fraction containing the soluble proteins is retained for further purification of the desired compound. If the product is secreted from the *C. glutamicum*

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cells, then the cells are removed from the culture by low-speed centrifugation, and the supernate fraction is retained for further purification.

The supernatant fraction from either purification method is subjected to chromatography with a suitable resin, in which the desired molecule is either retained on a chromatography resin while many of the impurities in the sample are not, or where the impurities are retained by the resin while the sample is not. Such chromatography steps may be repeated as necessary, using the same or different chromatography resins. One of ordinary skill in the art would be well-versed in the selection of appropriate chromatography resins and in their most efficacious application for a particular molecule to be purified. The purified product may be concentrated by filtration or ultrafiltration, and stored at a temperature at which the stability of the product is maximized.

There are a wide array of purification methods known to the art and the preceding method of purification is not meant to be limiting. Such purification techniques are described, for example, in Bailey, J.E. & Ollis, D.F. Biochemical Engineering Fundamentals, McGraw-Hill: New York (1986).

The identity and purity of the isolated compounds may be assessed by techniques standard in the art. These include high-performance liquid chromatography (HPLC), spectroscopic methods, staining methods, thin layer chromatography, NIRS, enzymatic assay, or microbiologically. Such analysis methods are reviewed in: Patek *et al.* (1994) *Appl. Environ. Microbiol.* 60: 133-140; Malakhova *et al.* (1996) *Biotekhnologiya* 11: 27-32; and Schmidt *et al.* (1998) *Bioprocess Engineer*. 19: 67-70. Ulmann's Encyclopedia of Industrial Chemistry, (1996) vol. A27, VCH: Weinheim, p. 89-90, p. 521-540, p. 540-547, p. 539-300, 575-381 and p. 581-587; Michal, G. (1999) Biochemical Pathways: An Atlas of Biochemistry and Molecular Biology, John Wiley and Sons; Fallon, A. *et al.* (1987) Applications of HPLC in Biochemistry in: Laboratory Techniques in Biochemistry and Molecular Biology, vol. 17.

Example 11: Analysis of the Gene Sequences of the Invention

The comparison of sequences and determination of percent homology between two sequences are art-known techniques, and can be accomplished using a mathematical algorithm, such as the algorithm of Karlin and Altschul (1990) *Proc. Natl. Acad. Sci.* USA 87:2264-68, modified as in Karlin and Altschul (1993) *Proc. Natl. Acad. Sci.* USA

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90:5873-77. Such an algorithm is incorporated into the NBLAST and XBLAST programs (version 2.0) of Altschul, *et al.* (1990) *J. Mol. Biol.* 215:403-10. BLAST nucleotide searches can be performed with the NBLAST program, score = 100, wordlength = 12 to obtain nucleotide sequences homologous to MP nucleic acid molecules of the invention. BLAST protein searches can be performed with the XBLAST program, score = 50, wordlength = 3 to obtain amino acid sequences homologous to MP protein molecules of the invention. To obtain gapped alignments for comparison purposes, Gapped BLAST can be utilized as described in Altschul *et al.*, (1997) *Nucleic Acids Res.* 25(17):3389-3402. When utilizing BLAST and Gapped BLAST programs, one of ordinary skill in the art will know how to optimize the parameters of the program (*e.g.*, XBLAST and NBLAST) for the specific sequence being analyzed.

Another example of a mathematical algorithm utilized for the comparison of sequences is the algorithm of Meyers and Miller ((1988) *Comput. Appl. Biosci.* 4: 11-17). Such an algorithm is incorporated into the ALIGN program (version 2.0) which is part of the GCG sequence alignment software package. When utilizing the ALIGN program for comparing amino acid sequences, a PAM120 weight residue table, a gap length penalty of 12, and a gap penalty of 4 can be used. Additional algorithms for sequence analysis are known in the art, and include ADVANCE and ADAM. described in Torelli and Robotti (1994) *Comput. Appl. Biosci.* 10:3-5; and FASTA, described in Pearson and Lipman (1988) *P.N.A.S.* 85:2444-8.

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The percent homology between two amino acid sequences can also be accomplished using the GAP program in the GCG software package (available at http://www.gcg.com), using either a Blosum 62 matrix or a PAM250 matrix, and a gap weight of 12, 10, 8, 6, or 4 and a length weight of 2, 3, or 4. The percent homology between two nucleic acid sequences can be accomplished using the GAP program in the GCG software package, using standard parameters, such as a gap weight of 50 and a length weight of 3.

A comparative analysis of the gene sequences of the invention with those present in Genbank has been performed using techniques known in the art (see, e.g., Bexevanis and Ouellette, eds. (1998) Bioinformatics: A Practical Guide to the Analysis of Genes and Proteins. John Wiley and Sons: New York). The gene sequences of the invention

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were compared to genes present in Genbank in a three-step process. In a first step, a BLASTN analysis (e.g., a local alignment analysis) was performed for each of the sequences of the invention against the nucleotide sequences present in Genbank, and the top 500 hits were retained for further analysis. A subsequent FASTA search (e.g., a combined local and global alignment analysis, in which limited regions of the sequences are aligned) was performed on these 500 hits. Each gene sequence of the invention was subsequently globally aligned to each of the top three FASTA hits, using the GAP program in the GCG software package (using standard parameters). In order to obtain correct results, the length of the sequences extracted from Genbank were adjusted to the length of the query sequences by methods well-known in the art. The results of this analysis are set forth in Table 4. The resulting data is identical to that which would have been obtained had a GAP (global) analysis alone been performed on each of the genes of the invention in comparison with each of the references in Genbank, but required significantly reduced computational time as compared to such a database-wide GAP (global) analysis. Sequences of the invention for which no alignments above the cutoff values were obtained are indicated on Table 4 by the absence of alignment information. It will further be understood by one of ordinary skill in the art that the GAP alignment homology percentages set forth in Table 4 under the heading "% homology (GAP)" are listed in the European numerical format, wherein a ',' represents a decimal point. For example, a value of "40,345" in this column represents "40.345%".

Example 12: Construction and Operation of DNA Microarrays

The sequences of the invention may additionally be used in the construction and application of DNA microarrays (the design, methodology, and uses of DNA arrays are well known in the art, and are described, for example, in Schena, M. et al. (1995) Science 270: 467-470; Wodicka, L. et al. (1997) Nature Biotechnology 15: 1359-1367; DeSaizieu, A. et al. (1998) Nature Biotechnology 16: 45-48; and DeRisi, J.L. et al. (1997) Science 278: 680-686).

DNA microarrays are solid or flexible supports consisting of nitrocellulose, nylon, glass, silicone, or other materials. Nucleic acid molecules may be attached to the surface in an ordered manner. After appropriate labeling, other nucleic acids or nucleic acid mixtures can be hybridized to the immobilized nucleic acid molecules, and the label

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may be used to monitor and measure the individual signal intensities of the hybridized molecules at defined regions. This methodology allows the simultaneous quantification of the relative or absolute amount of all or selected nucleic acids in the applied nucleic acid sample or mixture. DNA microarrays, therefore, permit an analysis of the expression of multiple (as many as 6800 or more) nucleic acids in parallel (see, *e.g.*, Schena, M. (1996) *BioEssays* 18(5): 427-431).

The sequences of the invention may be used to design oligonucleotide primers which are able to amplify defined regions of one or more *C. glutamicum* genes by a nucleic acid amplification reaction such as the polymerase chain reaction. The choice and design of the 5' or 3' oligonucleotide primers or of appropriate linkers allows the covalent attachment of the resulting PCR products to the surface of a support medium described above (and also described, for example, Schena, M. *et al.* (1995) *Science* 270: 467-470).

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Nucleic acid microarrays may also be constructed by *in situ* oligonucleotide synthesis as described by Wodicka, L. *et al.* (1997) *Nature Biotechnology* 15: 1359-1367. By photolithographic methods, precisely defined regions of the matrix are exposed to light. Protective groups which are photolabile are thereby activated and undergo nucleotide addition, whereas regions that are masked from light do not undergo any modification. Subsequent cycles of protection and light activation permit the synthesis of different oligonucleotides at defined positions. Small, defined regions of the genes of the invention may be synthesized on microarrays by solid phase oligonucleotide synthesis.

The nucleic acid molecules of the invention present in a sample or mixture of nucleotides may be hybridized to the microarrays. These nucleic acid molecules can be labeled according to standard methods. In brief, nucleic acid molecules (e.g., mRNA molecules or DNA molecules) are labeled by the incorporation of isotopically or fluorescently labeled nucleotides, e.g., during reverse transcription or DNA synthesis. Hybridization of labeled nucleic acids to microarrays is described (e.g., in Schena, M. et al. (1995) supra; Wodicka, L. et al. (1997), supra; and DeSaizieu A. et al. (1998), supra). The detection and quantification of the hybridized molecule are tailored to the specific incorporated label. Radioactive labels can be detected, for example, as

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described in Schena, M. et al. (1995) supra) and fluorescent labels may be detected, for example, by the method of Shalon et al. (1996) Genome Research 6: 639-645).

The application of the sequences of the invention to DNA microarray technology, as described above, permits comparative analyses of different strains of *C. glutamicum* or other Corynebacteria. For example, studies of inter-strain variations based on individual transcript profiles and the identification of genes that are important for specific and/or desired strain properties such as pathogenicity, productivity and stress tolerance are facilitated by nucleic acid array methodologies. Also, comparisons of the profile of expression of genes of the invention during the course of a fermentation reaction are possible using nucleic acid array technology.

Example 13: Analysis of the Dynamics of Cellular Protein Populations (Proteomics)

The gencs, compositions, and methods of the invention may be applied to study the interactions and dynamics of populations of proteins, termed 'proteomics'. Protein populations of interest include, but are not limited to, the total protein population of *C. glutamicum* (*e.g.*, in comparison with the protein populations of other organisms), those proteins which are active under specific environmental or metabolic conditions (*e.g.*, during fermentation, at high or low temperature, or at high or low pH), or those proteins which are active during specific phases of growth and development.

Protein populations can be analyzed by various well-known techniques, such as gel electrophoresis. Cellular proteins may be obtained, for example, by lysis or outraction, and may be separated from one another using a variety of electrophoretic techniques. Sodium dodecyl sulfate polyacrylamide gel electrophoresis (SDS-PAGE) separates proteins largely on the basis of their molecular weight. Isoelectric focusing polyacrylamide gel electrophoresis (IEF-PAGE) separates proteins by their isoelectric point (which reflects not only the amino acid sequence but also posttranslational modifications of the protein). Another, more preferred method of protein analysis is the consecutive combination of both IEF-PAGE and SDS-PAGE, known as 2-D-gel electrophoresis (described, for example, in Hermann et al. (1998) Electrophoresis 19: 3217-3221; Fountoulakis et al. (1998) Electrophoresis 19: 1193-1202; Langen et al. (1997) Electrophoresis 18: 1184-1192; Antelmann et al. (1997) Electrophoresis 18:

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1451-1463). Other separation techniques may also be utilized for protein separation, such as capillary gel electrophoresis; such techniques are well known in the art.

Proteins separated by these methodologies can be visualized by standard techniques, such as by staining or labeling. Suitable stains are known in the art, and include Coomassie Brilliant Blue, silver stain, or fluorescent dyes such as Sypro Ruby (Molecular Probes). The inclusion of radioactively labeled amino acids or other protein precursors (*e.g.*, ³⁵S-methionine, ³⁵S-cysteine, ¹⁴C-labelled amino acids, ¹⁵N-amino acids, ¹⁵NO₃ or ¹⁵NH₄⁺ or ¹³C-labelled amino acids) in the medium of *C. glutamicum* permits the labeling of proteins from these cells prior to their separation. Similarly, fluorescent labels may be employed. These labeled proteins can be extracted, isolated and separated according to the previously described techniques.

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Proteins visualized by these techniques can be further analyzed by measuring the amount of dye or label used. The amount of a given protein can be determined quantitatively using, for example, optical methods and can be compared to the amount of other proteins in the same gel or in other gels. Comparisons of proteins on gels can be made, for example, by optical comparison, by spectroscopy, by image scanning and analysis of gels, or through the use of photographic films and screens. Such techniques are well-known in the art.

To determine the identity of any given protein, direct sequencing or other standard techniques may be employed. For example, N- and/or C-terminal amino acid sequencing (such as Edman degradation) may be used, as may mass spectrometry (in particular MALDI or ESI techniques (see, e.g., Langen et al. (1997) Electrophoresis 18: 1184-1192)). The protein sequences provided herein can be used for the identification of *C. glutamicum* proteins by these techniques.

The information obtained by these methods can be used to compare patterns of protein presence, activity, or modification between different samples from various biological conditions (e.g., different organisms, time points of fermentation, media conditions, or different biotopes, among others). Data obtained from such experiments alone, or in combination with other techniques, can be used for various applications, such as to compare the behavior of various organisms in a given (e.g., metabolic) situation, to increase the productivity of strains which produce fine chemicals or to increase the efficiency of the production of fine chemicals.

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Equivalents

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Those of ordinary skill in the art will recognize, or will be able to ascertain using no more than routine experimentation, many equivalents to the specific embodiments of the invention described herein. Such equivalents are intended to be encompassed by the following claims.

What is claimed:

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- An isolated nucleic acid molecule from Corynebacterium glutamicum encoding a
 metabolic pathway protein, or a portion thereof, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.
 - 2. The isolated nucleic acid molecule of claim 1, wherein said metabolic pathway protein is selected from the group consisting of proteins involved in the metabolism of an amino acid, a vitamin, a cofactor, a nutraceutical, a nucleotide, a nucleoside, or trehalose.
- An isolated Corynebacterium glutamicum nucleic acid molecule selected from the group consisting of those sequences set forth as odd-numbered SEQ ID NOs of the
 Sequence Listing, or a portion thereof, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.
- 4. An isolated nucleic acid molecule which encodes a polypeptide sequence selected from the group consisting of those sequences set forth as even-numbered SEQ ID
 NOs of the Sequence Listing, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.
 - 5. An isolated nucleic acid molecule which encodes a naturally occurring allelic variant of a polypeptide selected from the group of amino acid sequences consisting of those sequences set forth as even-numbered SEQ ID NOs of the Sequence Listing, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.
- An isolated nucleic acid molecule comprising a nucleotide sequence which is at least
 50% homologous to a nucleotide sequence selected from the group consisting of those sequences set forth as odd-numbered SEQ ID NOs of the Sequence Listing, or

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a portion thereof, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.

- 7. An isolated nucleic acid molecule comprising a fragment of at least 15 nucleotides of a nucleic acid comprising a nucleotide sequence selected from the group consisting of those sequences set forth as odd-numbered SEQ ID NOs of the Sequence Listing, provided that the nucleic acid molecule does not consist of any of the F-designated genes set forth in Table 1.
- 10 8. An isolated nucleic acid molecule which hybridizes to the nucleic acid molecule of any one of claims 1-7 under stringent conditions.
 - 9. An isolated nucleic acid molecule comprising the nucleic acid molecule of any one of claims 1-8 or a portion thereof and a nucleotide sequence encoding a heterologous polypeptide.
 - 10. A vector comprising the nucleic acid molecule of any one of claims 1-9.
 - 11. The vector of claim 10, which is an expression vector.

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- 12. A host cell transfected with the expression vector of claim 11.
- 13. The host cell of claim 12, wherein said cell is a microorganism.
- 25 14. The host cell of claim 13, wherein said cell belongs to the genus *Corynebacterium* or *Brevibacterium*.
 - 15. The host cell of claim 12, wherein the expression of said nucleic acid molecule results in the modulation in production of a fine chemical from said cell.
 - 16. The host cell of claim 15, wherein said fine chemical is selected from the group consisting of: organic acids, nonproteinogenic amino acids, purine and pyrimidine

bases, nucleosides, nucleotides, lipids, saturated and unsaturated fatty acids, diols, carbohydrates, aromatic compounds, vitamins, cofactors, polyketides, and enzymes.

- 17. A method of producing a polypeptide comprising culturing the host cell of claim 12
 in an appropriate culture medium to, thereby, produce the polypeptide.
 - 18. An isolated metabolic pathway polypeptide from *Corynebacterium glutamicum*, or a portion thereof.
- 10 19. The protein of claim 18, wherein said polypeptide is selected from the group of metabolic pathway proteins which participate in the metabolism of an amino acid, a vitamin, a cofactor, a nutraccutical, a nucleotide, a nucleoside, or trehalose.
- 20. An isolated polypeptide comprising an amino acid sequence selected from the group
 consisting of those sequences set forth as even-numbered SEQ ID NOs of the
 Sequence Listing, provided that the amino acid sequence is not encoded by any of
 the F-designated genes set forth in Table 1.
- 21. An isolated polypeptide comprising a naturally occurring allelic variant of a polypeptide comprising an amino acid sequence selected from the group consisting of those sequences set forth as even-numbered SEQ ID NOs of the Sequence Listing, or a portion thereof, provided that the amino acid sequence is not encoded by any of the F-designated genes set forth in Table 1.
- 25 22. The isolated polypeptide of any of claims 18-21, further comprising heterologous amino acid sequences.
 - 23. An isolated polypeptide which is encoded by a nucleic acid molecule comprising a nucleotide sequence which is at least 50% homologous to a nucleic acid selected from the group consisting of those sequences set forth as odd-numbered SEQ ID NOs of the Sequence Listing, provided that the nucleic acid molecule does not consist of any of the F-designated nucleic acid molecules set forth in Table 1.

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- 24. An isolated polypeptide comprising an amino acid sequence which is at least 50% homologous to an amino acid sequence selected from the group consisting of those sequences set forth as even-numbered SEQ ID NOs of the Sequence Listing, provided that the amino acid sequence is not encoded by any of the F-designated genes set forth in Table 1.
- 25. A method for producing a fine chemical, comprising culturing a cell containing a vector of claim 12 such that the fine chemical is produced.
- 26. The method of claim 25, wherein said method further comprises the step of recovering the fine chemical from said culture.
- 27. The method of claim 25, wherein said method further comprises the step of
 transfecting said cell with the vector of claim 11 to result in a cell containing said vector.
 - 28. The method of claim 25, wherein said cell belongs to the genus *Corynebacterium* or *Brevibacterium*.
 - 29. The method of claim 25, wherein said cell is selected from the group consisting of: Corynebacterium glutamicum, Corynebacterium herculis, Corynebacterium, lilium, Corynebacterium acetoacidophilum, Corynebacterium acetogiulamicum,
- Corynebacterium acetophilum, Corynebacterium ammoniagenes, Corynebacterium fujiokense, Corynebacterium nitrilophilus, Brevibacterium ammoniagenes, Brevibacterium butanicum, Brevibacterium divaricatum, Brevibacterium flavum, Brevibacterium healii, Brevibacterium ketoglutamicum, Brevibacterium ketosoreductum, Brevibacterium lactofermentum, Brevibacterium linens, Brevibacterium paraffinolyticum, and those strains set forth in Table 3.
 - 30. The method of claim 25, wherein expression of the nucleic acid molecule from said vector results in modulation of production of said fine chemical.

- 31. The method of claim 25, wherein said fine chemical is selected from the group consisting of: organic acids, nonproteinogenic amino acids, purine and pyrimidine bases, nucleosides, nucleotides, lipids, saturated and unsaturated fatty acids, diols, carbohydrates, aromatic compounds, vitamins, cofactors, polyketides, and enzymes.
- 32. The method of claim 25, wherein said fine chemical is an amino acid.

- 33. The method of claim 32, wherein said amino acid is drawn from the group consisting of: lysine, glutamate, glutamine, alanine, aspartate, glycine, serine, threonine, methionine, cysteine, valine, leucine, isoleucine, arginine, proline, histidine, tyrosine, phenylalanine, and tryptophan.
- 34. A method for producing a fine chemical, comprising culturing a cell whose genomic
 DNA has been altered by the inclusion of a nucleic acid molecule of any one of claims 1-9.
- 35. A method for diagnosing the presence or activity of Corynebacterium diphtheriae in a subject, comprising detecting the presence of one or more of SEQ ID NOs 1
 through 1156 of the Sequence Listing in the subject, provided that the sequences are not or are not encoded by any of the F-designated sequences set forth in Table 1, thereby diagnosing the presence or activity of Corynebacterium diphtheriae in the subject.
- 25 36. A host cell comprising a nucleic acid molecule selected from the group consisting of the nucleic acid molecules set forth as odd-numbered SEQ ID NOs of the Sequence Listing, wherein the nucleic acid molecule is disrupted.
- 37. A host cell comprising a nucleic acid molecule selected from the group consisting of
 30 the nucleic acid molecules set forth as odd-numbered SEQ ID NOs in the Sequence
 Listing, wherein the nucleic acid molecule comprises one or more nucleic acid

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modifications from the sequence set forth as odd-numbered SEQ ID NOs of the Sequence Listing s.

38. A host cell comprising a nucleic acid molecule selected from the group consisting of the nucleic acid molecules set forth as odd-numbered SEQ ID NOs of the Sequence Listing, wherein the regulatory region of the nucleic acid molecule is modified relative to the wild-type regulatory region of the molecule.

SEQUENCE LISTING

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Asn Lys His Gly Ile Leu Phe Ile Ala Asp Glu Val Met Val Gly Phe 260 265 270

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75

7.0

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Ala Val Ala Asn Leu Leu Ala Thr Lys His Arg Gly Pro Asp Met Pro 50 55 60

Val Pro Val Leu Val Gly Ser Trp Asp Thr Ile Gln Gly Leu Val His 65 70 75 80

Ser Tyr Ser Ala Gln Ala Lys Ala Leu Val Glu Ala Phe Trp Pro Gly 90 Gly Leu Ser Ile Ile Val Pro Gln Ala Pro Ser Leu Pro Trp Asn Leu 105 Gly Asp Thr Arg Gly Thr Val Met Leu Arg Met Pro Leu His Pro Val 120 Ala Ile Glu Leu Leu Arg Gln Thr Gly Pro Met Ala Val Ser Ser Ala 135 Asn Ile Ser Gly His Thr Pro Pro Thr Thr Val Leu Glu Ala Arg Gln 155 150 Gln Leu Asn Gln Asn Val Ala Val Tyr Leu Asp Gly Gly Glu Cys Ala 170 165 Leu Ala Thr Pro Ser Thr Ile Val Asp Ile Ser Gly Pro Ala Pro Lys 185 Ile Leu Arg Glu Gly Ala Ile Ser Ala Glu Arg Val Gly Glu Val Leu 205 200 Gly Val Ser Ala Glu Ser Leu Arg 210 <210> 13 <211> 1026 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1003) <223> RXC00657 <400> 13 gtgcggatcg ggtatccgcg ctacacttag aggtgttaga gatcatgagt ttccacgaac 60 tgtaacgcag gattcaccaa tcaatgaaag gtcgaccgac atg agc act gaa gac Met Ser Thr Glu Asp 1 att gtc gtc gta gca gta gat ggc tcg gac gcc tca aaa caa gct gtt 163 Ile Val Val Val Ala Val Asp Gly Ser Asp Ala Ser Lys Gln Ala Val 15 10 cgg tgg gct gca aat acc gcc aac aaa cgt ggc att cca ctt cgc ttg Arg Trp Ala Ala Asn Thr Ala Asn Lys Arg Gly Ile Pro Leu Arg Leu 25 get tee age tae ace atg cet cag tte etc tae gea gag gga atg gtt Ala Ser Ser Tyr Thr Met Pro Gln Phe Leu Tyr Ala Glu Gly Met Val 45 40 cca cca caa gag ctt ttc gat gac ctc cag gcc gaa gcc ctg gaa aag Pro Pro Gln Glu Leu Phe Asp Asp Leu Gln Ala Glu Ala Leu Glu Lys 60 55

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						gca Ala										739
-						atg Met 220									_	787
					Val	acc Thr				Ile					Arg	835
230					233					240					245	
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Ala Glu Gly Met Val Pro Pro Gln Glu Leu Phe Asp Asp Leu Gln Ala 50 60

Glu Ala Leu Glu Lys Ile Asn Glu Ala Arg Asp Ile Ala His Glu Val 65 70 75 80

Ala Pro Glu Ile Lys Ile Gly His Thr Ile Ala Glu Gly Ser Pro Ile 85 90 95

Asp Met Leu Leu Glu Met Ser Pro Asp Ala Thr Met Ile Val Met Gly 100 105 110

Ser Arg Gly Leu Gly Gly Leu Ser Gly Met Val Met Gly Ser Val Ser 115 120 125

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Asp Ser Ala Val Asn Glu Asp Ser Lys Tyr Gly Pro Val Val Val Gly 145 150 150

Val Asp Gly Ser Glu Val Ser Gln Gln Ala Thr Glu Tyr Ala Phe Ala 165 170 175

Glu Ala Glu Ala Arg Gly Ala Glu Leu Val Ala Val His Thr Trp Met 180 185 190

Asp Met Gln Val Gln Ala Ser Leu Ala Gly Leu Ala Ala Ala Gln Gln 195 200 205

Gln Trp Asp Glu Val Glu Arg Gln Gln Thr Asp Met Leu Ile Glu Arg 210 215 220

Leu Ala Pro Leu Val Glu Lys Tyr Pro Ser Val Thr Val Lys Lys Ile 225 230 235 240

Ile Thr Arg Asp Arg Pro Val Arg Ala Leu Ala Glu Ala Ser Glu Asn 245 250 255

Ala Gln Leu Leu Val Val Gly Ser His Gly Arg Gly Gly Phe Lys Gly 260 265 270

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Asn Le	3	5				4	0				4	5			
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gct ttg 1363 Ala Leu															
1363	Val	Leu	Ser 410	Glu	Phe	Ala	Gly	Ala 415	Ala	Thr	Glu	Leu	Thr 420	Gly	

BNSDOCID - WO 0100843A2 1_>

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gtg gca gct gtc cat gat ttg aag cac aat ccg gaa tct gcg gca acg 1459 Val Ala Ala Val His Asp Leu Lys His Asp Pro Glu Ser Ala Ala Thr

cga atg aaa acg aac agc gag cag gtc tat acc cac gac gtc aac gtg 1507

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<213> Corynebacterium glutamicum

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Cys Trp Val Gly Trp Pro Gly Thr Val Asp Val Ala Pro Glu Pro Phe 50 60

Arg Thr Asp Thr Gly Val Leu Leu His Pro Val Val Leu Thr Ala Ser 65 70 75 80

Asp Tyr Glu Gly Phe Tyr Glu Gly Phe Ser Asn Ala Thr Leu Trp Pro 85 90 95

Leu Phe His Asp Leu Ile Val Thr Pro Val Tyr Asn Thr Asp Trp Trp 100 105 110

His Ala Phe Arg Glu Val Asn Leu Lys Phe Ala Glu Ala Val Ser Gln 115 120 125

Val Ala Ala His Gly Ala Thr Val Trp Val Gln Asp Tyr Gln Leu Leu 130 135 140

Leu Val Pro Gly Ile Leu Arg Gln Met Arg Pro Asp Leu Lys Ile Gly 145 150 155 160

Phe Phe Leu His Ile Pro Phe Pro Ser Pro Asp Leu Phe Arg Gln Leu 165 170 175

Pro Trp Arg Glu Glu Ile Val Arg Gly Met Leu Gly Ala Asp Leu Val 180 185 190

Gly Phe His Leu Val Gln Asn Ala Glu Asn Phe Leu Ala Leu Thr Gln 200 Gln Val Ala Gly Thr Ala Gly Ser His Val Gly Gln Pro Asp Thr Leu 210 215 Gln Val Ser Gly Glu Ala Leu Val Arg Glu Ile Gly Ala His Val Glu Thr Ala Asp Gly Arg Arg Val Ser Val Gly Ala Phe Pro Ile Ser Ile 245 250 Asp Val Glu Met Phe Gly Glu Ala Ser Lys Ser Ala Val Leu Asp Leu 260 265 Leu Lys Thr Leu Asp Glu Pro Glu Thr Val Phe Leu Gly Val Asp Arg 280 285 Leu Asp Tyr Thr Lys Gly Ile Leu Gln Arg Leu Leu Ala Phe Glu Glu Leu Leu Glu Ser Gly Ala Leu Glu Ala Asp Lys Ala Val Leu Leu Gln 310 315 Val Ala Thr Pro Ser Arg Glu Arg Ile Asp His Tyr Arg Val Ser Arg 330 Ser Gln Val Glu Ala Val Gly Arg Ile Asn Gly Arg Phe Gly Arg 340 345 Met Gly Arg Pro Val Val His Tyr Leu His Arg Ser Leu Ser Lys Asn 360 Asp Leu Gln Val Leu Tyr Thr Ala Ala Asp Val Met Leu Val Thr Pro 375 380 Phe Lys Asp Gly Met Asn Leu Val Ala Lys Glu Phe Val Ala Asn His 390 395 Arg Asp Gly Thr Gly Ala Leu Val Leu Ser Glu Phe Ala Gly Ala Ala 410 405 415 Thr Glu Leu Thr Gly Ala Tyr Leu Cys Asn Pro Phe Asp Val Glu Ser 420 425 Ile Lys Arg Gln Met Val Ala Ala Val His Asp Leu Lys His Asn Pro 435 440 445 Glu Ser Ala Ala Thr Arg Met Lys Thr Asn Ser Glu Gln Val Tyr Thr 455 His Asp Val Asn Val Trp Ala Asn Ser Phe Leu Asp Cys Leu Ala Gln 470 475 Ser Gly Glu Asn Ser

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caa aac Gln Asn															739
gcc ggg Ala Gly 215															787
gca ttg Ala Leu 230															835
cga gtt Arg Val															883
ggg gag Gly Glu															931
gag ccg Glu Pro															979
ggc att	ttg	cag	cgc	ctg	ctt	gcg	ttt	gag	gaa	ctg	ctg	gaa	tcc	ggc	
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Ala Tyr Leu Cys Asn Pro Phe Asp Val Glu Ser Ile Lys Arg Gln Met 425 430 435

gtg gca gct gtc cat gat ttg aag cac aat ccg gaa tct gcg gca acg 1459

Val Ala Ala Val His Asp Leu Lys His Asn Pro Glu Ser Ala Ala Thr 440 445 450

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Arg Met Lys Thr Asn Ser Glu Gln Val Tyr Thr His Asp Val Asn Val 455 460 465

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Gly Gly Leu Val Thr Gly Leu Ser Pro Val Leu Glu Gln His Arg Gly 35 40 45

Cys Trp Val Gly Trp Pro Gly Thr Val Asp Val Ala Pro Glu Pro Phe 50 60

Arg Thr Asp Thr Gly Val Leu Leu His Pro Val Val Leu Thr Ala Ser 65 70 75 80

Asp Tyr Glu Gly Phe Tyr Glu Gly Phe Ser Asn Ala Thr Leu Trp Pro 85 90 95

Leu Phe His Asp Leu Ile Val Thr Pro Val Tyr Asn Thr Asp Trp Trp 100 105 110

His Ala Phe Arg Glu Val Asn Leu Lys Phe Ala Glu Ala Val Ser Gln
115 120 125

Val Ala Ala His Gly Ala Thr Val Trp Val Gln Asp Tyr Gln Leu Leu 130 135 140

Leu Val Pro Gly Ile Leu Arg Gln Met Arg Pro Asp Leu Lys Ile Gly 145 150 155 160

Phe Phe Leu His Ile Pro Phe Pro Ser Pro Asp Leu Phe Arg Gln Leu 170 Pro Trp Arg Glu Glu Ile Val Arg Gly Met Leu Gly Ala Asp Leu Val Gly Phe His Leu Val Gln Asn Ala Glu Asn Phe Leu Ala Leu Thr Gln Gln Val Ala Gly Thr Ala Gly Ser His Val Gly Gln Pro Asp Thr Leu 215 Gln Val Ser Gly Glu Ala Leu Val Arg Glu Ile Gly Ala His Val Glu 230 Thr Ala Asp Gly Arg Arg Val Ser Val Gly Ala Phe Pro Ile Ser Ile 250 Asp Val Glu Met Phe Gly Glu Ala Ser Lys Ser Ala Val Leu Asp Leu Leu Lys Thr Leu Asp Glu Pro Glu Thr Val Phe Leu Gly Val Asp Arg 280 Leu Asp Tyr Thr Lys Gly Ile Leu Gln Arg Leu Leu Ala Phe Glu Glu Leu Leu Glu Ser Gly Ala Leu Glu Ala Asp Lys Ala Val Leu Leu Gln Val Ala Thr Pro Ser Arg Glu Arg Ile Asp His Tyr Arg Val Ser Arg 330 Ser Gln Val Glu Glu Ala Val Gly Arg Ile Asn Gly Arg Phe Gly Arg Met Gly Arg Pro Val Val His Tyr Leu His Arg Ser Leu Ser Lys Asn 360 Asp Leu Gln Val Leu Tyr Thr Ala Ala Asp Val Met Leu Val Thr Pro 375 Phe Lys Asp Gly Met Asn Leu Val Ala Lys Glu Phe Val Ala Asn His 395 390 Arg Asp Gly Thr Gly Ala Leu Val Leu Ser Glu Phe Ala Gly Ala Ala 410 Thr Glu Leu Thr Gly Ala Tyr Leu Cys Asn Pro Phe Asp Val Glu Ser 425 Ile Lys Arg Gln Met Val Ala Ala Val His Asp Leu Lys His Asn Pro Glu Ser Ala Ala Thr Arg Met Lys Thr Asn Ser Glu Gln Val Tyr Thr 455 His Asp Val Asn Val Trp Ala Asn Ser Phe Leu Asp Cys Leu Ala Gln 475 470 465

Ser Gly

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185 180 190 atg age aaa tat eet eag gea gte teg ett gat ttg egt gaa ttt gea 624 Met Ser Lys Tyr Pro Gln Ala Val Ser Leu Asp Leu Arg Glu Phe Ala 200 195 205 gga cac acc cct cga gag atg tcg ggc ggg cag ctg ttc cct acc att 672 Gly His Thr Pro Arg Glu Met Ser Gly Gly Gln Leu Phe Pro Thr Ile 210 215 gct gaa cgg gag tgg att gtc act tta gcc cct cac gga ttc ttc tgg Ala Glu Arg Glu Trp Ile Val Thr Leu Ala Pro His Gly Phe Phe Trp 225 766 ttt gat ctc acc gcc gat gaa aag gac gat atg gaa tgagcattgg Phe Asp Leu Thr Ala Asp Glu Lys Asp Asp Met Glu 779 ccaacacatc atc <210> 22 <211> 252 <212> PRT <213> Corynebacterium glutamicum <400> 22 Thr Ala Gln Trp Gly Ile Phe Leu Arg Asn His Asp Glu Leu Thr Leu Glu Met Val Ser Asp Glu Glu Arg Ser Tyr Met Tyr Ser Gln Phe Ala Ser Glu Pro Arg Met Arg Ala Asn Val Gly Ile Arg Arg Arg Leu Ser Pro Leu Leu Glu Gly Asp Arg Asn Gln Leu Glu Leu His Gly Leu Leu Leu Ser Leu Pro Gly Ser Pro Val Leu Tyr Tyr Gly Asp Glu Ile 70 75 met Gly Asp Asn lie Trp Leu His Asp Arg Asp Gly Val Arg Thr Pro Met Gln Trp Ser Asn Asp Arg Asn Gly Gly Phe Ser Lys Ala Asp Pro Glu Arg Leu Tyr Leu Pro Ala Ile Gln Asn Asp Gln Tyr Gly Tyr Ala Gln Val Asn Val Glu Ser Gln Leu Asn Arg Glu Asn Ser Leu Leu 135 Arg Trp Leu Arg Asn Gln Ile Leu Ile Arg Lys Gln Tyr Arg Ala Phe 150 Gly Ala Gly Thr Tyr Arg Glu Val Ser Ser Thr Asn Glu Ser Val Leu 170 Thr Phe Leu Arg Glu His Lys Gly Gln Thr Ile Leu Cys Val Asn Asn

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Gly	His 210	Thr	Pro	Arg	Glu	Met 215	Ser	Gly	Gly	Gln	Leu 220	Phe	Pro	Thr	Ile	
Ala 225	Glu .	Arg	Glu	Trp	Ile 230	Val	Thr	Leu	Ala	Pro 235	His	Gly	Phe	Phe	Trp 240	
Phe	Asp	Leu	Thr	Ala 245	Asp	Glu	Lys	Asp	Asp 250	Met	Glu					
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	_	_		-		_	_							gac Asp		595
														tcc Ser 180		643
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														gaa Glu		739
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														gtc Val		835
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<211> 334

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<212> PRT

<213> Corynebacterium glutamicum

<400> 24

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Thr Ser Pro Leu Asn Ser Gln Pro Ser Ala Asp His His Pro Asp His 20 25 30

Ala Ala Arg Pro Val Leu Asp Ala His Gly Leu Ile Val Glu His Glu 35 40 45

Ser Glu Glu Phe Pro Val Pro Ala Pro Ala Pro Gly Glu Gln Pro Trp 50 55 60

Glu Lys Lys Asn Arg Glu Trp Tyr Lys Asp Ala Val Phe Tyr Glu Val 65 70 75 80

Leu Val Arg Ala Phe Tyr Asp Pro Glu Gly Asn Gly Val Gly Ser Leu
85 90 95

Lys Gly Leu Thr Glu Lys Leu Asp Tyr Ile Gln Trp Leu Gly Val Asp 100 105 110

Cys Ile Trp Ile Pro Pro Phe Tyr Asp Ser Pro Leu Arg Asp Gly Gly
115 120 125

Tyr Asp Ile Arg Asn Phe Arg Glu Ile Leu Pro Glu Phe Gly Thr Val 130 135 140

Asp Asp Phe Val Glu Leu Val Asp His Ala His Arg Arg Gly Leu Arg 145 150 155 160

Val Ile Thr Asp Leu Val Met Asn His Thr Ser Asp Gln His Ala Trp 165 170 175

Phe Gln Glu Ser Arg Arg Asp Pro Thr Gly Pro Tyr Gly Asp Phe Tyr 180 185 190

Val Trp Ser Asp Asp Pro Thr Leu Tyr Asn Glu Ala Arg Ile Ile Phe 195 200 205

Val Asp Thr Glu Glu Ser Asn Trp Thr Tyr Asp Pro Val Arg Gly Gln 210 215 220

Tyr Phe Trp His Arg Phe Phe Ser His Gln Pro Asp Leu Asn Tyr Asp 225 230 235 240

Asn Pro Ala Val Gln Glu Ala Met Leu Asp Val Leu Arg Phe Trp Leu 245 250 255

Asp Leu Gly Leu Asp Gly Phe Arg Leu Asp Ala Val Pro Tyr Leu Phe 260 265 270

Glu Arg Glu Gly Thr Asn Gly Glu Asn Leu Lys Glu Thr His Asp Phe 275 280 285

Leu Lys Leu Cys Arg Ser Val Ile Glu Lys Glu Tyr Pro Gly Arg Ile 290 295 300

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Gly 150	Arg	Gly	Gly	Ser	Asp 155	Thr	Thr	Ala	Val	Ala 160	Leu	Ala	Ala	Ala	Leu 165	
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gct Ala	gac Asp	ccg Pro	cgc Arg 185	atc Ile	gtt Val	cct Pro	aat Asn	gca Ala 190	cag Gln	aag Lys	ctg Leu	gaa Glu	aag Lys 195	ctc Leu	agc Ser	691
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Leu	Lys	Lys	Leu	Gln 330		Gln	Gly	Asn	Trp 335		Asn	Val	Leu	Туr 340	Asp	
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Pro	o Gly	7 Val		Ala	a Glu	Phe	Met 365		ı Ala	a Leu	a Arg	Asp 370		. Asr	n Val	

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Arg Glu Asp Asp Leu Asp Ala Ala Ala Arg Ala Leu His Glu Gln Phe 390 395 400 405

cag ctg ggc ggc gaa gac gaa gcc gtc gtt tat gca ggc acc gga cgc 1363

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Gly Asn Asp Val Val Val Cys Ser Ala Met Gly Asp Thr Thr Asp \$35\$ \$40\$ \$45\$

Glu Leu Leu Glu Leu Ala Ala Ala Val Asn Pro Val Pro Pro Ala Arg 50 55 60

Glu Met Asp Met Leu Leu Thr Ala Gly Glu Arg Ile Ser Asn Ala Leu 65 70 75 80

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Gly Ser Gln Ala Gly Val Leu Thr Thr Glu Arg His Gly Asn Ala Arg 100 105 110

Ile Val Asp Val Thr Pro Gly Arg Val Arg Glu Ala Leu Asp Glu Gly 115 120 125

Lys Ile Cys Ile Val Ala Gly Phe Gln Gly Val Asn Lys Glu Thr Arg 130 135 140

Asp Val Thr Thr Leu Gly Arg Gly Gly Ser Asp Thr Thr Ala Val Ala 145 150 155 160

Leu Ala Ala Ala Leu Asn Ala Asp Val Cys Glu Ile Tyr Ser Asp Val 165 170 175

Asp Gly Val Tyr Thr Ala Asp Pro Arg Ile Val Pro Asn Ala Gln Lys
180 185 190

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										gag Glu			787
										ctc Leu			835

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Val Asn Pro Ser Asp Lys Asp Ser Leu Val Lys Gly Ile Ile Ala Asn 120 Pro Asn Cys Thr Thr Met Ala Ala Met Pro Val Leu Lys Pro Leu His 135 140 Asp Ala Ala Gly Leu Val Lys Leu His Val Ser Ser Tyr Gln Ala Val Ser Gly Ser Gly Leu Ala Gly Val Glu Thr Leu Ala Lys Gln Val Ala 170 Ala Val Gly Asp His Asn Val Glu Phe Val His Asp Gly Gln Ala Ala 185 Asp Ala Gly Asp Val Gly Pro Tyr Val Ser Pro Ile Ala Tyr Asn Val 200 Leu Pro Phe Ala Gly Asn Leu Val Asp Asp Gly Thr Phe Glu Thr Asp 215 Glu Glu Gln Lys Leu Arg Asn Glu Ser Arg Lys Ile Leu Gly Leu Pro 235 230 Asp Leu Lys Val Ser Gly Thr Cys Val Arg Val Pro Val Phe Thr Gly 250 His Thr Leu Thr Ile His Ala Glu Phe Asp Lys Ala Ile Thr Val Asp Gin Ala Gln Glu Ile Leu Gly Ala Ala Ser Gly Val Lys Leu Val Asp 280 Val Pro Thr Pro Leu Ala Ala Gly Ile Asp Glu Ser Leu Val Gly Arg Ile Arg Gln Asp Ser Thr Val Asp Asp Asn Arg Gly Leu Val Leu 310 Val Val Ser Gly Asp Asn Leu Arg Lys Gly Ala Ala Leu Asn Thr Ile 325 330

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Asp	) Lev	ı Asr	n Glu 265	ı Ala	ı Ile	Glu	His	270	. Val	l Gli	ı Thr	. Leu	275	i Leu	gac Asp	931
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150

145

Ala Leu Leu Gly Glu Pro Thr Gly Gly Trp Ile Glu Ala Gly Cys Gln 170 Gly Asn Leu Arg Ile Lys Val Thr Ala His Gly Val Arg Ala His Ser 185 Ala Arg Ser Trp Leu Gly Asp Asn Ala Met His Lys Leu Ser Pro Ile Ile Ser Lys Val Ala Ala Tyr Lys Ala Ala Glu Val Asn Ile Asp Gly 215 Leu Thr Tyr Arg Glu Gly Leu Asn Ile Val Phe Cys Glu Ser Gly Val Ala Asn Asn Val Ile Pro Asp Leu Ala Trp Met Asn Leu Asn Phe Arg 250 245 Phe Ala Pro Asn Arg Asp Leu Asn Glu Ala Ile Glu His Val Val Glu Thr Leu Glu Leu Asp Gly Gln Asp Gly Ile Glu Trp Ala Val Glu Asp 280 Gly Ala Gly Gly Ala Leu Pro Gly Leu Gly Gln Gln Val Thr Ser Gly Leu Ile Asp Ala Val Gly Arg Glu Lys Ile Arg Ala Lys Phe Gly Trp 315 310 Thr Asp Val Ser Arg Phe Ser Ala Met Gly Ile Pro Ala Leu Asn Phe Gly Ala Gly Asp Pro Ser Phe Ala His Lys Arg Asp Glu Gln Cys Pro 345 Val Glu Gln Ile Thr Asp Val Ala Ala Ile Leu Lys Gln Tyr Leu Ser 360 Glu <210> 33 <211> 1059 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1036) <223> RXA00044 <400> 33 attacctcag ccttccaagc tgatgatgca ttacttaaaa actgcagaca cttgaaaaac 60 ttctcacccg cactcgttcc ctcaacccac aaggagcacc atg gct tcc gca act Met Ala Ser Ala Thr ttc acc ggc gtg atc cca ccc gta atg acc cca ctc cac gcc gac ggc

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H					cgc Arg												355
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					gcc Ala												451
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age gae gaa gaa act get ege att eac gee att gtt gat gaa tte etg 1027

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Ser Gly Glu Ala Ala Phe Leu Thr Arg Ala Gln Arg Lys Leu Ala Leu 50 55 60

Thr Thr Ile Ile Glu His Thr Ala Gly Arg Val Pro Val Thr Ala Gly 65 70 75 80

Val Ile Glu Thr Thr Thr Ala Arg Val Ile Glu Leu Val Glu Asp Ala 85 90 95

Leu Glu Ala Gly Ala Glu Gly Leu Val Ala Thr Ala Pro Phe Tyr Thr  $100 \hspace{1.5cm} 105 \hspace{1.5cm} 110$ 

Arg Thr His Asp Val Glu Ile Glu Glu His Phe Arg Lys Ile His Ala 115 120 125

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His Ser Asn Leu Asn Pro Val Met Leu Leu Thr Leu Ala Lys Asp Gly 145 150 155 160

Val Leu Ala Gly Thr Lys Asp Ser Ser Gly Asp Gly Ala Ile Arg 165 170 175

Ser Leu Ile Glu Ala Arg Asp Asp Ala Gly Leu Thr Glu Gln Phe Lys

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Pro	His 290	Gln	Ser	Leu	Ser	Asp 295	Glu	Glu	Thr	Ala	Arg 300	Ile	His	Ala	Ile	
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Thi	r Ile	e Val	l Ala 20	_	a Val	l Ası	n Glu	ı Ser 25		) Ası	) Lev	ı Glu	ı Lev 30	ı Val	Ala	
Gli	ı Ile	e Gl	y Val	l Asp	As <u>r</u>	) Ası	, Le	ı Sei	c Lei	ı Lei	ı Val	l Ası	Ası	ı Gly	/ Ala	

35 40 45 Glu Val Val Asp Phe Thr Thr Pro Asn Ala Val Met Gly Asn Leu 55 Glu Phe Cys Ile Asn Asn Gly Ile Ser Ala Val Val Gly Thr Thr Gly 75 Phe Asp Asp Ala Arg Leu Glu Gln Val Arg Asp Trp Leu Glu Gly Lys Asp Asn Val Gly Val Leu Ile Ala Pro Asn Phe Ala Ile Ser Ala Val 105 Leu Thr Met Val Phe Ser Lys Gln Ala Ala Arg Phe Phe Glu Ser Ala 120 Glu Val Ile Glu Leu His His Pro Asn Lys Leu Asp Ala Pro Ser Gly 135 Thr Ala Ile His Thr Ala Gln Gly Ile Ala Ala Arg Lys Glu Ala 150 Gly Met Asp Ala Gln Pro Asp Ala Thr Glu Gln Ala Leu Glu Gly Ser 170 Arg Gly Ala Ser Val Asp Gly Ile Pro Val His Ala Val Arg Met Ser 185 Gly Met Val Ala His Glu Gln Val Ile Phe Gly Thr Gln Gly Gln Thr Leu Thr Ile Lys Gln Asp Ser Tyr Asp Arg Asn Ser Phe Ala Pro Gly 215 Val Leu Val Gly Val Arg Asn Ile Ala Gln His Pro Gly Leu Val Val 230 Gly Leu Glu His Tyr Leu Gly Leu 245

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agcagcgggt ggaatttttt aaaaggagcg tttaaaggct gtg gcc gaa caa gtt  $$\,$  115  $$\,$  Val Ala Glu Gln Val  $$\,$  1  $\,$  5

aaa ttg age gtg gag ttg ata geg tge agt tet ttt act eea eee get 163 Lys Leu Ser Val Glu Leu Ile Ala Cys Ser Ser Phe Thr Pro Pro Ala

				10					15					20		
gat Asp	gtt Val	gag Glu	tgg Trp 25	tca Ser	act Thr	gat Asp	gtt Val	gag Glu 30	ggc Gly	gcg Ala	gaa Glu	gca Ala	ctc Leu 35	gtc Val	gag Glu	211
ttt Phe	gcg Ala	ggt Gly 40	cgt Arg	gcc Ala	tgc Cys	tac Tyr	gaa Glu 45	act Thr	ttt Phe	gat Asp	aag Lys	ccg Pro 50	aac Asn	cct Pro	cga Arg	259
act Thr	gct Ala 55	tcc Ser	aat Asn	gct Ala	gcg Ala	tat Tyr 60	ctg Leu	cgc Arg	cac His	atc Ile	atg Met 65	gaa Glu	gtg Val	ggg Gly	cac His	307
act Thr 70	gct Ala	ttg Leu	ctt Leu	gag Glu	cat His 75	gcc Ala	aat Asn	gcc Ala	acg Thr	atg Met 80	tat Tyr	atc Ile	cga Arg	ggc Gly	att Ile 85	355
tct Ser	cgg Arg	tcc Ser	gcg Ala	acc Thr 90	cat His	gaa Glu	ttg Leu	gtc Val	cga Arg 95	cac His	cgc Arg	cat His	ttt Phe	tcc Ser 100	ttc Phe	403
tct Ser	caa Gln	ctg Leu	tct Ser 105	cag Gln	cgt Arg	ttc Phe	gtg Val	cac His 110	agc Ser	gga Gly	gaa Glu	tcg Ser	gaa Glu 115	gta Val	gtg Val	451
gtg Val	ccc Pro	act Thr 120	ctc Leu	atc Ile	gat Asp	gaa Glu	gat Asp 125	ccg Pro	cag Gln	ttg Leu	cgt Arg	gaa Glu 130	ctt Leu	ttc Phe	atg Met	499
cac His	gcc Ala 135	Met	gat Asp	gag Glu	tct Ser	cgg Arg 140	Phe	gct Ala	ttc Phe	aat Asn	gag Glu 145	Leu	ctt Leu	aat Asn	gcg Ala	547
ctg Leu 150	Glu	gaa Glu	aaa Lys	ctt Leu	ggc Gly 155	Asp	gaa Glu	. ccg . Pro	aat Asn	gca Ala 160	ı Leu	tta Leu	agg Arg	aaa Lys	aag Lys 165	595
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aga Arg	a ato	gtg Val	g gtg Val 185	Ser	gga Gly	aac / Asr	tto Phe	cgc Arg 190	y Thi	tgg Tr	g ago p Aro	g cat g His	tto Phe 195	; ITE	ggc Gly	691
at Me	g cga t Arg	a gco g Ala 200	a Sei	gaa Glu	a cat ı His	gca Ala	a gad a Asp 205	va.	gaa l Glu	a ato ı Ile	c cgo e Aro	g gaa g Glu 210	ıval	a gcg . Ala	g gta a Val	739
ga Gl	a Lgi u Cy: 21	s Le	a aga u Arg	a aaq g Lys	g cto s Lev	g cag u Gli 220	n Val	a gca l Ala	agco aAla	g cca a Pro	a act o Th: 22!	r va.	t tto l Phe	ggt Gly	t gat y Asp	787
tt Ph 23	e Gl	g at u Il	t ga e Gl	a act u Thi	t ttg r Le 23	u Ala	a ga a Asj	c gg p Gl	a tc y Se	g ca r Gl 24	n Me	g gc. t Al	a aca	a ago r Se:	c ccg r Pro 245	835
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       Met Thr Thr Ala Ser Ala Thr Gly Ile Ala Thr Leu Thr Ser
acc ggc gac gtc ctg gac gtg tgg tat cca gaa atc ggg tcc acc gac
                                                                  158
Thr Gly Asp Val Leu Asp Val Trp Tyr Pro Glu Ile Gly Ser Thr Asp
15
                     20
                                         25
cag tee geg ete aca eet eta gaa gge gte gat gaa gat ega aac gte
                                                                  206
Gln Ser Ala Leu Thr Pro Leu Glu Gly Val Asp Glu Asp Arg Asn Val
                 35
                                                                  254
acc cgc aaa atc gtg acg aca act atc gac acc gac gca gcc ccc acc
Thr Arg Lys Ile Val Thr Thr Thr Ile Asp Thr Asp Ala Ala Pro Thr
             5.0
                                 55
gac acc tac gat gca tgg ctg cgc ctt cac ctc ctc tcc cac cgc gtt
                                                                  302
Asp Thr Tyr Asp Ala Trp Leu Arg Leu His Leu Leu Ser His Arg Val
         65
                            7.0
ttc cgc cct cac acc atc aac cta gac ggc att ttc ggc ctc ctc aac
                                                                  350
Phe Arg Pro His Thr Ile Asn Leu Asp Gly Ile Phe Gly Leu Leu Asn
     80
                         85
aat gtc gtg tgg acc aac ttc gga ccg tgc gca gtt gac ggt ttc gca
                                                                  398
Asn Val Val Trp Thr Asn Phe Gly Pro Cys Ala Val Asp Gly Phe Ala
95
                    100
                                        105
ctc acc cgc gcg cgc ctg tca cgc cga ggc caa gtt acg gtt tat agc
Leu Thr Arg Ala Arg Leu Ser Arg Gly Gln Val Thr Val Tyr Ser
                115
                                    120
gtc gac aag ttc cca cgc atg gtc gac tat gtg gtt ccc tcg ggc gtg
                                                                  494
Val Asp Lys Phe Pro Arg Met Val Asp Tyr Val Val Pro Ser Gly Val
           130
                                135
cgc atc ggt gac gcc gac cgc gtc cga ctt ggc gcg tac ctg gca gat
Arg Ile Gly Asp Ala Asp Arg Val Arg Leu Gly Ala Tyr Leu Ala Asp
                            150
       145
gge ace ace gtg atg cat gag gge tte gtg aac tte aac get gge acg
                                                                  590
Gly Thr Thr Val Met His Glu Gly Pne Val Asn Phe Asn Ala Gly Thr
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Leu Gly Ala Ser Met Val
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Ala Leu Thr Pro Leu Glu Gly Val Asp Glu Asp Arg Asn Val Thr Arg 35 40 45

Lys Ile Val Thr Thr Thr Ile Asp Thr Asp Ala Ala Pro Thr Asp Thr 50 60

Tyr Asp Ala Trp Leu Arg Leu His Leu Leu Ser His Arg Val Phe Arg 65 70 75 80

Pro His Thr Ile Asn Leu Asp Gly Ile Phe Gly Leu Leu Asn Asn Val 85 90 95

Val Trp Thr Asn Phe Gly Pro Cys Ala Val Asp Gly Phe Ala Leu Thr
100 105 110

Arg Ala Arg Leu Ser Arg Arg Gly Gln Val Thr Val Tyr Ser Val Asp 115 120 125

Lys Phe Pro Arg Met Val Asp Tyr Val Val Pro Ser Gly Val Arg Ile 130 140

Gly Asp Ala Asp Arg Val Arg Leu Gly Ala Tyr Leu Ala Asp Gly Thr 145 150 155 160

Thr Val Met His Glu Gly Phe Val Asn Phe Asn Ala Gly Thr Leu Gly 165 170 175

Ala Ser Met Val 180

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ggtcctgatg aaagagatgt ccctgaatca tcatctaagt atg cat ctc ggt aag 115
Met His Leu Gly Lys
1

ctc gac cag gac agt gcc acc aca att ttg gag gat tac aag aac atg  $\,$  163 Leu Asp Gln Asp Ser Ala Thr Thr Ile Leu Glu Asp Tyr Lys Asn Met  $\,$  10  $\,$  15  $\,$  20

			-				gtg Val									211
_	-	_			-		cag Gln 45		_							259
	_	_		-			gac Asp									307
_	_		_	_		-	gac Asp	-		-		-		_	-	355
							cct Pro									403
							tac Tyr									451
							gcc Ala 125									499
_							cca Pro									547
							gag Glu									595
							tcc Ser									643
gtt Val	caa Gln	aag Lys	gca Ala 185	gtc Val	cag Gln	tac Tyr	acc Thr	ctc Leu 190	cca Pro	tcc Ser	gaa Glu	gac Asp	gcc Ala 195	ctg Leu	gaa Glu	691
							ggc Gly 205									739
	_		_			-	gcc Ala	-		-	-			-		787
							cct Pro									835
•	-				_		gca Ala									883
atg	cca	cac	ggt	ggc	cac	gtg	att	acc	acc	ggc	gac	acc	ggt	ggc	ttc	931

Met Pro His Gly Gly His Val Ile Thr Thr Gly Asp Thr Gly Gly Phe 265 270 979 aac cac acc gtg gaa tac atc ctc aag ctg gac cga aac cca gat ttc Asn His Thr Val Glu Tyr Ile Leu Lys Leu Asp Arg Asn Pro Asp Phe 285 acc get tee tea cag ate get tte ggt ege gea get eac ege atg aag 1027 Thr Ala Ser Ser Gln Ile Ala Phe Gly Arg Ala Ala His Arg Met Lys cag cag ggc caa agc gga gct ttc acc gtc ctc gaa gtt gct cca tac Gln Gln Gly Gln Ser Gly Ala Phe Thr Val Leu Glu Val Ala Pro Tyr 315 310 ctg ctc tcc cca gag aac ttg gac gat ctg atc gca cgc gac gtc 1120 Leu Leu Ser Pro Glu Asn Leu Asp Asp Leu Ile Ala Arg Asp Val 330 335 taatttagct cgaggggcaa gga 1143 <210> 42 <211> 340 <212> PRT <213> Corynebacterium glutamicum <400> 42 Met His Leu Gly Lys Leu Asp Gln Asp Ser Ala Thr Thr Ile Leu Glu Asp Tyr Lys Asn Met Thr Asn Ile Arg Val Ala Ile Val Gly Tyr Gly Asn Leu Gly Arg Ser Val Glu Lys Leu Ile Ala Lys Gln Pro Asp Met <u>Asp Leu Val Gly Ile Phe Ser Arg Arg Ala Thr Leu Asp Thr Lys Thr</u> Pro Val Phe Asp Val Ala Asp Val Asp Lys His Ala Asp Asp Val Asp Val Leu Phe Leu Cys Met Gly Ser Ala Thr Asp Ile Pro Glu Gln Ala Pro Lys Phe Ala Gln Phe Ala Cys Thr Val Asp Thr Tyr Asp Asn His 105 Arg Asp Ile Pro Arg His Arg Gln Val Met Asn Glu Ala Ala Thr Ala 120 Ala Gly Asn Val Ala Leu Val Ser Thr Gly Trp Asp Pro Gly Met Phe 135 Ser Ile Asn Arg Val Tyr Ala Ala Ala Val Leu Ala Glu His Gln Gln 155

His Thr Fhe Trp Gly Pro Gly Leu Ser Gln Gly His Ser Asp Ala Leu

165 170

Arg Arg Ile Pro Gly Val Gln Lys Ala Val Gln Tyr Thr Leu Pro Ser Glu Asp Ala Leu Glu Lys Ala Arg Arg Gly Glu Ala Gly Asp Leu Thr 200 Gly Lys Gln Thr His Lys Arg Gln Cys Phe Val Val Ala Asp Ala Ala Asp His Glu Arg Ile Glu Asn Asp Ile Arg Thr Met Pro Asp Tyr Phe 235 230 Val Gly Tyr Glu Val Glu Val Asn Phe Ile Asp Glu Ala Thr Phe Asp Ser Glu His Thr Gly Met Pro His Gly Gly His Val Ile Thr Thr Gly 265 Asp Thr Gly Gly Phe Asn His Thr Val Glu Tyr Ile Leu Lys Leu Asp 280 Arg Asn Pro Asp Phe Thr Ala Ser Ser Gln Ile Ala Phe Gly Arg Ala 295 Ala His Arg Met Lys Gln Gln Gly Gln Ser Gly Ala Phe Thr Val Leu 310 305 Glu Val Ala Pro Tyr Leu Leu Ser Pro Glu Asn Leu Asp Asp Leu Ile 330 325 Ala Arg Asp Val 340 <210> 43 <211> 958 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(958) <223> FRXA00352 <400> 43 aatagatcag cgcatccgtg gtggaaccaa aaggctcaac aatacgaaac gttcgctttc 60 agtcctgatg aaagagatgt ccctgaatca tcatctaagt atg cat ctc ggt aag Met His Leu Gly Lys ctc gac cag gac agt gcc acc aca att ttg gag gat tac aag aac atg Leu Asp Gln Asp Ser Ala Thr Thr Ile Leu Glu Asp Tyr Lys Asn Met acc aac atc cgc gta gct atc gtg ggc tac gga aac ctg gga cgc agc Thr Asn Ile Arg Val Ala Ile Val Gly Tyr Gly Asn Leu Gly Arg Ser

			25					30					35			
gtc g Val G		_			-	_	_		-	_	_		_			259
ttc to																307
gcc g Ala A 70																355
atg g Met G																403
ttc g Phe A		_		-	_											451
cac c His A	rg	_	~	_		-	_	-		_				-		499
ctg g Leu V 1																547
tac g Tyr A 150																595
cca g Pro G	_	_		-				_	_	_	_	_				643
gtt c Val G		_	_	_	_						_		_	_	_	691
aag g Lys A																739
aag c Lys A 2																787
gaa a Glu A 230		_				-		-			-			-	-	835
gaa g Glu V	•				-	_	_			_						883
atg c Met P																931

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958

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PCT/IB00/00923 WO 01/00843

260 270 265 Asp Thr Gly Gly Phe Asn His Thr Val Glu Tyr Ile Leu Lys 275 280 <210> 45 <211> 1400 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (1)..(1377) <223> RXA00972 <400> 45 cct gca cct ggt tgg cgt ttc cgc acc gga gaa gat gta aca atg gct 48 Pro Ala Pro Gly Trp Arg Phe Arg Thr Gly Glu Asp Val Thr Met Ala 10 aca gtt gaa aat ttc aat gaa ctt ccc gca cac gta tgg cca cgc aat 96 Thr Val Glu Asn Phe Asn Glu Leu Pro Ala His Val Trp Pro Arg Asn 25 20 gcc gtg cgc caa gaa gac ggc gtt gtc acc gtc ggt gtg cct ctg 144 Ala Val Arg Gln Glu Asp Gly Val Val Thr Val Ala Gly Val Pro Leu 35 cet gae etc get gaa gaa tac gga ace eea etg tte gta gte gae gag 192 Pro Asp Leu Ala Glu Glu Tyr Gly Thr Pro Leu Phe Val Val Asp Glu gac gat ttc cgt tcc cgc tgt cgc gac atg gct acc gca ttc ggt gga 240 Asp Asp Phe Arg Ser Arg Cys Arg Asp Met Ala Thr Ala Phe Gly Gly 288 cca ggc aat gtg cac tac gca tct aaa gcg ttc ctg acc aag acc att Pro Gly Asn Val His Tyr Ala Ser Lys Ala Phe Leu Thr Lys Thr Ile 90 gea egt tgg gtt gat gaa gag ggg etg gea etg gae att gea tee ate 336 Ala Arg Trp Val Asp Glu Glu Gly Leu Ala Leu Asp Ile Ala Ser Ile 100 105 aac gaa ctg ggc att gcc ctg gcc gct ttc ccc gcc agc cgt atc 384 Asn Glu Leu Gly Ile Ala Leu Ala Gly Phe Pro Ala Ser Arg Ile 115 120 acc gcg cac ggc aac aac aaa ggc gta gag ttc ctg cgc gcg ttg gtt 432 Thr Ala His Gly Asn Asn Lys Gly Val Glu Phe Leu Arg Ala Leu Val 130 135 caa aac ggt gtg gga cac gtg gtg ctg gac tcc gca cag gaa cta gaa 480 Gln Asn Gly Val Gly His Val Val Leu Asp Ser Ala Gln Glu Leu Glu 145 150 155 528 ctg ttg gat tac gtt gcc gct ggt gaa ggc aag att cag gac gtg ttg Leu Leu Asp Tyr Val Ala Ala Gly Glu Gly Lys Ile Gln Asp Val Leu 165

170

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act Thr	agc Ser	cac His 195	gaa Glu	gac Asp	cag Gln	aag Lys	ttc Phe 200	gga Gly	ttc Phe	tcc Ser	ctg Leu	gca Ala 205	tcc Ser	ggt Gly	tcc Ser	624
gca Ala	ttc Phe 210	gaa Glu	gca Ala	gca Ala	aaa Lys	gcc Ala 215	gcc Ala	aac Asn	aac Asn	gca Ala	gaa Glu 220	aac Asn	ctg Leu	aac Asn	ctg Leu	672
	ggc Gly															720
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agc Ser	gaa Glu	ctg Leu	ggc Gly 260	gtt Val	gcc Ala	ctt Leu	cct Pro	gaa Glu 265	ctg Leu	gat Asp	ctc Leu	ggt Gly	ggc Gly 270	gga Gly	tac Tyr	816
ggc Gly	att Ile	gcc Ala 275	tat Tyr	acc Thr	gca Ala	gct Ala	gaa Glu 280	gaa Glu	cca Pro	ctc Leu	aac Asn	gtc Val 285	gca Ala	gaa Glu	gtt Val	864
gcc Ala	tcc Ser 290	gac Asp	ctg Leu	ctc Leu	acc Thr	gca Ala 295	gtc Val	gga Gly	aaa Lys	atg Met	gca Ala 300	gcg Ala	gaa Glu	cta Leu	ggc Gly	912
atc Ile 305	gac Asp	gca Ala	cca Pro	acc Thr	gtg Val 310	ctt Leu	gtt Val	gag Glu	ccc Pro	ggc Gly 315	cgc Arg	gct Ala	atc Ile	gca Ala	ggc Gly 320	960
ccc 100	tcc g	acc	gtg	acc	atc	tac	gaa	gtc	ggc	acc	acc	aaa	gac	gtc	cac	
Pro	Ser	Thr	Val	Thr 325	Ile	Tyr	Glu	Val	Gly 330	Thr	Thr	Lys	Asp	Val 335	His	
gta 105	gac 6	gac	gac	aaa	acc	cgc	cgt	tac	atc	gcc	gtg	gac	gga	ggc	atg	
	Asp	Asp	Asp 340	Lys	Thr	Arg	Arg	Tyr 345	Ile	Ala	Val	Asp	Gly 350		Met	
tcc 110	gac 4	aac	atc	cgc	cca	gca	ctc	tac	ggg	tcc	gaa	tac	gac	gcc	cgc	
	Asp	Asn 355		Arg	Pro	Ala	Leu 360		Gly	Ser	Glu	Tyr 365		Ala	Arg	
gta 115		tcc	cgc	ttc	gcc	gaa	gga	gac	cca	gta	agc	acc	cgc	ato	gtg	
			Arg	Phe	Ala	Glu 375		Asp	Pro	Val	Ser 380		Arg	Ile	Val	
ggc 120		cac	tgc	gaa	tcc	ggc	gat	atc	ctg	atc	aac	gat	gaa	ato	tac	
	Ser	His	Cys	Glu	Ser 390		Asp	Ile	Leu	. Ile 395		Asp	Glu	Ile	Tyr 400	

cca tot gad atd add agd ggd gad ttd ott gda otd gda gdd add ggd Pro Ser Asp Ile Thr Ser Gly Asp Phe Leu Ala Leu Ala Ala Thr Gly gca tac tgc tac gcc atg agc tcc cgc tac aac gcc ttc aca cgg ccc 1296 Ala Tyr Cys Tyr Ala Met Ser Ser Arg Tyr Asn Ala Phe Thr Arg Pro 425 420 gcc gtc gtg tcc gtc cgc gct ggc agc tcc cgc ctc atg ctg cgc cgc 1344 Ala Val Val Ser Val Arg Ala Gly Ser Ser Arg Leu Met Leu Arg Arg 435 440 gaa acg ctc gac gac atc ctc tca cta gag gca taacgctttt cgacgcctga Glu Thr Leu Asp Asp Ile Leu Ser Leu Glu Ala CCC 1400 <210> 46 <211> 459 <212> PRT <213> Corynebacterium glutamicum <400> 46 Pro Ala Pro Gly Trp Arg Phe Arg Thr Gly Glu Asp Val Thr Met Ala Thr Val Glu Asn Phe Asn Glu Leu Pro Ala His Val Trp Pro Arg Asn 2.5 Ala Val Arg Gln Glu Asp Gly Val Val Thr Val Ala Gly Val Pro Leu Pro Asp Leu Ala Glu Glu Tyr Gly Thr Pro Leu Phe Val Val Asp Glu Asp Asp Phe Arg Ser Arg Cys Arg Asp Met Ala Thr Ala Phe Gly Gly Pro Gly Asn Val His Tyr Ala Ser Lys Ala Phe Leu Thr Lys Thr Ile Ala Arg Trp Val Asp Glu Glu Gly Leu Ala Leu Asp Ile Ala Ser Ile Asn Glu Leu Gly Ile Ala Leu Ala Ala Gly Phe Pro Ala Ser Arg Ile 120 Thr Ala His Gly Asn Asn Lys Gly Val Glu Phe Leu Arg Ala Leu Val Gln Asn Gly Val Gly His Val Val Leu Asp Ser Ala Gln Glu Leu Glu 150 155

Leu Leu Asp Tyr Val Ala Ala Gly Glu Gly Lys Ile Gln Asp Val Leu

165 170 Ile Arg Val Lys Pro Gly Ile Glu Ala His Thr His Glu Phe Ile Ala 185 Thr Ser His Glu Asp Gln Lys Phe Gly Phe Ser Leu Ala Ser Gly Ser 200 Ala Phe Glu Ala Ala Lys Ala Ala Asn Asn Ala Glu Asn Leu Asn Leu Val Gly Leu His Cys His Val Gly Ser Gln Val Phe Asp Ala Glu Gly 230 235 Phe Lys Leu Ala Ala Glu Arg Val Leu Gly Leu Tyr Ser Gln Ile His Ser Glu Leu Gly Val Ala Leu Pro Glu Leu Asp Leu Gly Gly Tyr 265 Gly Ile Ala Tyr Thr Ala Ala Glu Glu Pro Leu Asn Val Ala Glu Val 280 Ala Ser Asp Leu Leu Thr Ala Val Gly Lys Met Ala Ala Glu Leu Gly 295 Ile Asp Ala Pro Thr Val Leu Val Glu Pro Gly Arg Ala Ile Ala Gly 310 Pro Ser Thr Val Thr Ile Tyr Glu Val Gly Thr Thr Lys Asp Val His 330 Val Asp Asp Asp Lys Thr Arg Arg Tyr Ile Ala Val Asp Gly Gly Met Ser Asp Asn Ile Arg Pro Ala Leu Tyr Gly Ser Glu Tyr Asp Ala Arg 360 Val Val Ser Arg Phe Ala Glu Gly Asp Pro Val Ser Thr Arg Ile Val Gly Ser His Cys Glu Ser Gly Asp Ile Leu Ile Asn Asp Glu Ile Tyr 390 395 Pro Ser Asp Ile Thr Ser Gly Asp Phe Leu Ala Leu Ala Ala Thr Gly Ala Tyr Cys Tyr Ala Met Ser Ser Arg Tyr Asn Ala Phe Thr Arg Pro 425 Ala Val Val Ser Val Arg Ala Gly Ser Ser Arg Leu Met Leu Arg Arg 440

Glu Thr Leu Asp Asp Ile Leu Ser Leu Glu Ala

455

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<213> Corynebacterium glutamicum

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~ ~		-		_	_	_				_		atc Ile 210	-	~		739
												gat Asp				787
			_		_	-		-	-		~ ~	gcg Ala	_	_		835
												gaa Glu				883
									_		_	aat Asn				931
												cga Arg 290				979
		aaa	gcg	aat	aag	ggt	ctt	acc	ttc	gtt	gat	gcc	gtt	aaa	gac	
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		cgt	gga	gtc	cca	gga	gag	cgg	atc	att	cta	tcc	gca	gct	atc	
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	-	gac	aga	cta	ttg	gca	tta	gcg	atc	gaa	aat	ggc	gtg	atc	atc	
1171 Lys		Asp	Arg 345	Leu	Leu	Ala	Leu	Ala 350	Ile	Glu	Asn	Gly	Val 355	Ile	Ile	
tct 1219		gat	tcg	cgt	gat	gaa	tta	gat	cgc	att	tcg	gct	ttg	gtt	ggt	
		Asp 360	Ser	Arg	Asp	Glu	Leu 365	Asp	Arg	Ile	Ser	Ala 370	Leu	Val	Gly	
_	_	gtt	gca	cga	gtt	gcg	cct	aga	gta	gct	сса	gat	cct	gca	gtc	
1267 Asp		Val	Ala	Arg	Val	Ala 380	Pro	Arg	Val	Ala	Pro 385	Asp	Pro	Ala	Val	
		cca	act	aga	ttt	ggt	gag	cgt	gct	gca	gac	tgg	ggt	aat	cgg	
1315 Leu 390		Pro	Thr	Arg	Phe 395	Gly	Glu	Arg	Ala	Ala 400	Asp	Trp	Gly	Asn	Arg 405	
ctt 1363		gag	gtg	ata	ccc	ggc	gtg	gat	att	gtg	ggt	ctt	cac	gtt	cac	

Leu Thr Glu Val Ile Pro Gly Val Asp Ile Val Gly Leu His Val His 410 415 ctc cat ggc tat gct gca aaa gac cgt gct ctg gct ctg cag gaa tgt 1411 Leu His Gly Tyr Ala Ala Lys Asp Arg Ala Leu Ala Leu Gln Glu Cys 425 430 tgc caa ctc gtc gat tct ctc aga gaa tgc ggg cat tcc cca cag ttt 1459 Cys Gln Leu Val Asp Ser Leu Arg Glu Cys Gly His Ser Pro Gln Phe 445 att gac ctt gga ggg gtg cct atg agc tac att gaa tct gag gaa 1507 Ile Asp Leu Gly Gly Gly Val Pro Met Ser Tyr Ile Glu Ser Glu Glu 460 455 gat tgg atc cgt tat caa tcc gct aaa tct gcg act tca gcc ggg tat 1555 Asp Trp Ile Arg Tyr Gln Ser Ala Lys Ser Ala Thr Ser Ala Gly Tyr 475 480 470 gcc gaa tcc ttt acg tgg aaa gac gat ccg tta tct aat acg tac ccg 1603 Ala Glu Ser Phe Thr Trp Lys Asp Asp Pro Leu Ser Asn Thr Tyr Pro 490 495 ttc tat cag acc cca gtg cgc ggt aat tgg ttg aaa gac gtg ctt tct 1651 Phe Tyr Gln Thr Pro Val Arg Gly Asn Trp Leu Lys Asp Val Leu Ser 505 510 aag ggg gta gct cag atg ctc att gac cgg gga ttg cgg tta cac ata 1699 Lys Gly Val Ala Gln Met Leu Ile Asp Arg Gly Leu Arg Leu His Ile 520 525 530 gag cet ggt cga agt tta eta gat ggg tgt ggc gtc act ett gee gaa 1747 Glu Pro Gly Arg Ser Leu Leu Asp Gly Cys Gly Val Thr Leu Ala Glu 540 gtt gct ttt gtg aaa acc cga agt gac ggg ttg cct cta gtg gga ctg 1795 Val Ala Phe Val Lys Thr Arg Ser Asp Gly Leu Pro Leu Val Gly Leu 555 560 550 gct atg aac cga acg cag tgc cgg act aca tcc gat gat ttt ctc att 1843 Ala Met Asn Arg Thr Gln Cys Arg Thr Thr Ser Asp Asp Phe Leu Ile 570 575 gat ccc ctg cat atc act gac ggt gat gta ggc gag gaa atc gaa gca 1891 Asp Pro Leu His Ile Thr Asp Gly Asp Val Gly Glu Glu Ile Glu Ala 590 tat cta gtg ggt gcc tac tgc atc gaa gat gag ctg att tta cgc cgg Tyr Leu Val Gly Ala Tyr Cys Ile Glu Asp Glu Leu Ile Leu Arg Arg

600 605 610

cga atc cgc ttc ccg aga gga gtc aaa cca gga gat atc atc gga att 1987

Arg Ile Arg Phe Pro Arg Gly Val Lys Pro Gly Asp Ile Ile Gly Ile 615 620 625

cct aac acc gca gga tac ttc atg cat atc ttg gaa agt gca tcg cac 2035

Pro Asn Thr Ala Gly Tyr Phe Met His Ile Leu Glu Ser Ala Ser His 630 645

caa atc ccg ttg gcg aaa aat gta gtg tgg ccg gag ggg cag tta gac 2083

Gln Ile Pro Leu Ala Lys Asn Val Val Trp Pro Glu Gly Gln Leu Asp
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Asp Ile Asp Ala Asp 665

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<212> PRT

<213> Corynebacterium glutamicum

<400> 48

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Arg Thr Val Leu Lys Glu Val Ser Ser Gln Ile Gln Glu Arg Ala Gly  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Lys Lys Asp Glu Glu Trp Gly Met Gly Ala Thr Trp Arg Glu Leu Tyr 50 55 60

Pro Ser Ile Val Glu Arg Ala Ser Tyr Glu Gly Arg Asp Ser Leu Ile 65 70 75 80

Gly Phe Asp His Leu Ala Arg Glu Met Glu Arg Leu Ala Phe Gly Pro \$85\$ 90 95

Pro Ser Glu Ser Phe Glu Tyr Leu Gln Glu Leu Val Lys Ser Gly Val 100 105 110

Val Asp Ile Thr His Leu His Arg Gly Arg Glu Pro Leu Thr Asp Leu 115 120 125

Val Arg Glu Leu Glu Ile Thr Val Val Ile Asp Ala Val Leu Pro Pro 130 135 140

Pro Gly Val Val Pro Gly Thr Leu Val His Asn Leu Val Lys Glu Gly 145 150 155 160

Tyr Ala Arg Met Arg Pro Gly Thr Arg Gly Leu Asp Val Ala Ala Asp 165 170 175

Gly Thr Val Gln Gly Gln Arg His Leu Ala Ala Val Gly Arg Met Thr 180 Glu Asp Val Val Leu Gly Asn Asp Thr Leu Ser Arg Ser Leu His Asp 200 Ile Ile Pro Lys Trp Ala Arg Arg Val Ile Arg Asp Ala Ser Thr Tyr Pro Asp Arg Val His Gly Thr Pro Pro Leu Pro Ala Arg Leu Glu Pro 230 Trp Ala Glu Lys Leu Thr Ser Asp Pro Ala Thr Cys Arg His Leu Ile Glu Glu Phe Gly Ser Pro Val Asn Val Leu His Ser Gly Ser Met Pro Arg Asn Ile Asn Glu Leu Val Asp Ala Gly Ile Gln Met Gly Val Asp Thr Arg Ile Phe Phe Ala Arg Lys Ala Asn Lys Gly Leu Thr Phe Val Asp Ala Val Lys Asp Thr Gly His Gly Val Asp Val Ala Ser Glu Arg Glu Leu Ser Gln Val Leu Asn Arg Gly Val Pro Gly Glu Arg Ile Ile Leu Ser Ala Ala Ile Lys Pro Asp Arg Leu Leu Ala Leu Ala Ile Glu 345 Asn Gly Val Ile Ile Ser Val Asp Ser Arg Asp Glu Leu Asp Arg Ile 360 Ser Ala Leu Val Gly Asp Arg Val Ala Arg Val Ala Pro Arg Val Ala Pro Asp Pro Ala Val Leu Pro Pro Thr Arg Phe Gly Glu Arg Ala Ala Asp Trp Gly Asn Arg Leu Thr Glu Val Ile Pro Gly Val Asp Ile Val Gly Leu His Val His Leu His Gly Tyr Ala Ala Lys Asp Arg Ala Leu Ala Leu Gln Glu Cys Cys Gln Leu Val Asp Ser Leu Arg Glu Cys Gly His Ser Pro Gln Phe Ile Asp Leu Gly Gly Gly Val Pro Met Ser Tyr Ile Glu Ser Glu Glu Asp Trp Ile Arg Tyr Gln Ser Ala Lys Ser Ala 470 475 Thr Ser Ala Gly Tyr Ala Glu Ser Phe Thr Trp Lys Asp Asp Pro Leu

Ser Asn Thr Tyr Pro Phe Tyr Gln Thr Pro Val Arg Gly Asn Trp Leu 505 Lys Asp Val Leu Ser Lys Gly Val Ala Gln Met Leu Ile Asp Arg Gly 515 520 Leu Arg Leu His Ile Glu Pro Gly Arg Ser Leu Leu Asp Gly Cys Gly 535 Val Thr Leu Ala Glu Val Ala Phe Val Lys Thr Arg Ser Asp Gly Leu 550 Pro Leu Val Gly Leu Ala Met Asn Arg Thr Gln Cys Arg Thr Thr Ser Asp Asp Phe Leu Ile Asp Pro Leu His Ile Thr Asp Gly Asp Val Gly 585 Glu Glu Ile Glu Ala Tyr Leu Val Gly Ala Tyr Cys Ile Glu Asp Glu 595 600 Leu Ile Leu Arg Arg Ile Arg Phe Pro Arg Gly Val Lys Pro Gly 615 Asp Ile Ile Gly Ile Pro Asn Thr Ala Gly Tyr Phe Met His Ile Leu 630 Glu Ser Ala Ser His Gln Ile Pro Leu Ala Lys Asn Val Val Trp Pro 645 650 Glu Gly Gln Leu Asp Asp Ile Asp Ala Asp <210> 49 <211> 993 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(970) <223> RXA01393 <400> 49 caaaagcaga cctgtaatga agatttccat gatcaccatc gtgacctatg gaagtactta 60 agtaaaatga ttggttctta acatggttta atatagcttc atg aac ccc att caa Met Asn Pro Ile Gln ctg gac act ttg ctc tca atc att gat gaa ggc agc ttc gaa ggc gcc Leu Asp Thr Leu Leu Ser Ile Ile Asp Glu Gly Ser Phe Glu Gly Ala 10 the tha god off the att the ede tog gog gtg agt eag ege gtt aaa Ser Leu Ala Leu Ser Ile Ser Pro Ser Ala Val Ser Gln Arg Val Lys 25 gct ctc gag cat cac gtg ggt cga gtg ttg gta tcg cgc acc caa ccg Ala Leu Glu His His Val Gly Arg Val Leu Val Ser Arg Thr Gln Pro

		40					45			50				
	aaa Lys 55													307
	gtg Val													355
	gaa Glu													403
	ttt Phe													451
	acg Thr	_	-	_	-	-	-							499
	gga Gly 135													547
	tgt Cys													595
	ccc Pro													643
	gcg Ala													691
	gac Asp													739
	att Ile 215													787
	ggt Gly													835
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_	gcc Ala				_						tagt	taci	tc	980

tgaaaaggtt cag 993

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- <212> PRT
- <213> Corynebacterium glutamicum

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- Ser Phe Glu Gly Ala Ser Leu Ala Leu Ser Ile Ser Pro Ser Ala Val 20 25 30
- Ser Gln Arg Val Lys Ala Leu Glu His His Val Gly Arg Val Leu Val 35 40 45
- Ser Arg Thr Gln Pro Ala Lys Ala Thr Glu Ala Gly Glu Val Leu Val 50 60
- Gln Ala Ala Arg Lys Met Val Leu Leu Gln Ala Glu Thr Lys Ala Gln 65 70 75 80
- Leu Ser Gly Arg Leu Ala Glu Ile Pro Leu Thr Ile Ala Ile Asn Ala 85 90 95
- Asp Ser Leu Ser Thr Trp Phe Pro Pro Val Phe Asn Glu Val Ala Ser 100 105 110
- Trp Gly Gly Ala Thr Leu Thr Leu Arg Leu Glu Asp Glu Ala His Thr 115 120 125
- Leu Ser Leu Leu Arg Arg Gly Asp Val Leu Gly Ala Val Thr Arg Glu 130 140
- Ala Asn Pro Val Ala Gly Cys Glu Val Val Glu Leu Gly Thr Met Arg 145  $\phantom{\bigg|}$  150  $\phantom{\bigg|}$  155  $\phantom{\bigg|}$  160
- His Leu Ala Ile Ala Thr Pro Ser Leu Arg Asp Ala Tyr Met Val Asp 165 170 175
- Gly Lys Leu Asp Trp Ala Ala Met Pro Val Leu Arg Phe Gly Pro Lys  $180 \hspace{1cm} 185 \hspace{1cm} 190$
- Asp Val Leu Gln Asp Arg Asp Leu Asp Gly Arg Val Asp Gly Pro Val 195 200 205
- Gly Arg Arg Val Ser Ile Val Pro Ser Ala Glu Gly Phe Gly Glu 210 215 220
- Ala Ile Arg Arg Gly Leu Gly Trp Gly Leu Leu Pro Glu Thr Gln Ala 225 230 235 240
- Ala Pro Met Leu Lys Ala Gly Glu Val Ile Leu Leu Asp Glu Ile Pro 245 250 255
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Leu Val 150	Phe Gl	y Val	Val 155	Ser	Arg	Gly	Ile	Ser 160	Gln	Ala	Ala	Phe	Leu 165	
aca acg Thr Thr	_						_		_		-			643
atc ctt Ile Leu		a Phe				-								691
tta tgg Leu Trp		_					_				-			739
ggc atc Gly Ile 215														787
tcg gta Ser Val 230		_	-	_	-		_	-	_	_	_	-	-	835
acc gtg Thr Val														883
tcg ctg Ser Leu	-	e Gly		-										931
gat aat Asp Asn			_	_		_	_	_	_					979
gcc gca 1027	ttg at	t tcg	ttg	ggt	ctg	tgt	ctt	tcg	gtt	ctt	ggg	gcc	tat	
Ala Ala 295	Leu Il	e Ser	Leu	Gly 300	Leu	Cys	Leu	Ser	Val 305	Leu	Gly	Ala	Tyr	
gtg tcc 1075	tgg ca	g atg	ctc	tgc	gca	gaa	cca	ctg	gcg	ttg	atg	gca	atg	
Val Ser 310	Trp Gl	n Met	Leu 315	Cys	Ala	Glu	Pro	Leu 320	Ala	Leu	Met	Ala	Met 325	
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Asp Gly	Leu Il	e Pro 330		Lys	Ile	Gly	Ala 335	Ile	Asn	Ser	Arg	Gly 340	Ala	
gcc tgg 1171	atg gc	t cag	ctg	atc	tcc	acc	atc	gtg	att	cag	att	ttc	atc	
Ala Trp	Met Al 34		Leu	Ile	Ser	Thr 350	Ile	Val	Ile	Gln	Ile 355	Phe	Ile	
atc att 1219	ttc tt	c ctc	aac	gag	acc	acc	tac	gtc	tcc	atg	gtg	caa	ttg	
Ile Ile	Phe Ph 360	e Leu	Asn	Glu	Thr 365	Thr	Tyr	Val	Ser	Met 370	Val	Gln	Leu	

get ace aac cta tac ttg gtg cet tac etg tte tet gee ttt tat etg 1267 Ala Thr Asn Leu Tyr Leu Val Pro Tyr Leu Phe Ser Ala Phe Tyr Leu gtc atg ctg gca aca cgt gga aaa gga atc acc cac cca cat gcc ggc 1315 Val Met Leu Ala Thr Arg Gly Lys Gly Ile Thr His Pro His Ala Gly 395 aca cgt ttt gat gat tcc ggt cca gag ata tcc cgc cga gaa aac cgc Thr Arg Phe Asp Asp Ser Gly Pro Glu Ile Ser Arg Arg Glu Asn Arg 410 aaa cac ctc atc gtc ggt tta gta gca acg gtg tat tca gtg tgg ctg 1411 Lys His Leu Ile Val Gly Leu Val Ala Thr Val Tyr Ser Val Trp Leu ttt tac gct gca gaa ccg cag ttt gtc ctc ttc gga gcc atg gcg atg Phe Tyr Ala Ala Glu Pro Gln Phe Val Leu Phe Gly Ala Met Ala Met 445 ctt ccc ggc tta atc ccc tat gtg tgg aca agg att tat cgt ggc gaa Leu Pro Gly Leu Ile Pro Tyr Val Trp Thr Arg Ile Tyr Arg Gly Glu 460 cag gtg ttt aac cgc ttt gaa atc ggc gtg gtt gtt gtc ctg gtc gtt Gln Val Phe Asn Arg Phe Glu Ile Gly Val Val Val Leu Val Val 475 470 gct gcc agc gcg ggc gtt att ggt ttg gtc aac gga tca cta tcg ctt 1603 Ala Ala Ser Ala Gly Val Ile Gly Leu Val Asn Gly Ser Leu Ser Leu

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<210> 52

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		•		85		-		2	90	J		•		95	•
Tyr	Val	Gly	Phe 100	Ser	Ser	Ala	Trp	Gly 105	Tyr	Trp	Leu	Gly	Ser 110	Val	Ile
Ala	Gln	Val 115	Gly	Tyr	Ala	Thr	Leu 120	Phe	Phe	Ser	Thr	Leu 125	Gly	His	Tyr
Val	Pro 130	Leu	Phe	Ser	Gln	Asp 135	His	Pro	Phe	Val	Ser 140	Ala	Leu	Ala	Val
Ser 145	Ala	Leu	Thr	Trp	Leu 150	Val	Phe	Gly	Val	Val 155	Ser	Arg	Gly	Ile	Ser 160
Gln	Ala	Ala	Phe	Leu 165	Thr	Thr	Val	Thr	Thr 170	Val	Ala	Lys	Ile	Leu 175	Pro
Leu	Leu	Cys	Phe 180	Ile	Ile	Leu	Val	Ala 185	Phe	Leu	Gly	Phe	Ser 190	Trp	Glu
Lys	Phe	Thr 195	Val	Asp	Leu	Trp	Ala 200	Arg	Asp	Gly	Gly	Val 205	Gly	Ser	Ile
Phe	Asp 210	Gln	Val	Arg	Gly	Ile 215	Met	Val	Tyr	Thr	Val 220	Trp	Val	Phe	Ile
Gly 225	Ile	Glu	Gly	Ala	Ser 230	Val	Tyr	Ser	Arg	Gln 235	Ala	Arg	Ser	Arg	Ser 240
Asp	Val	Ser	Arg	Ala 245	Thr	Val	Ile	Gly	Phe 250	Val	Ala	Val	Leu	Leu 255	Leu
Leu	Val	Ser	Ile 260	Ser	Ser	Leu	Ser	Phe 265	Gly	Val	Leu	Thr	Gln 270	Gln	Glu
Leu	Ala	Ala 275	Leu	Pro	Asp	Asn	Ser 280	Met	Ala	Ser	Val	Leu 285	Glu	Ala	Val
Val	Gly 290	Pro	Trp	Gly	Ala	Ala 295	Leu	Ile	Ser	Leu	Gly 300	Leu	Cys	Leu	Ser
Val 305	Leu	Gly	Ala	Tyr	Val 310	Ser	Trp	Gln	Met	Leu 315	Cys	Ala	Glu	Pro	Leu 320
Ala	Leu	Met	Ala	Met 325	Asp	Gly	Leu	Ile	Pro 330	Ser	Lys	Ile	Gly	Ala 335	Ile
Asn	Ser	Arg	Gly 340	Ala	Ala	Trp	Met	Ala 345	Gln	Leu	Ile	Ser	Thr 350	Ile	Val
Ile	Gln	Ile 355	Phe	Ile	Ile	Ile	Phe 360	Phe	Leu	Asn	Glu	Thr 365	Thr	Tyr	Val
Ser	Met 370	Val	Gln	Leu	Ala	Thr 375	Asn	Leu	Tyr	Leu	Val 380	Pro	Tyr	Leu	Phe

Ser Ala Phe Tyr Leu Val Met Leu Ala Thr Arg Gly Lys Gly Ile Thr 390 His Pro His Ala Gly Thr Arg Phe Asp Asp Ser Gly Pro Glu Ile Ser 410 Arg Arg Glu Asn Arg Lys His Leu Ile Val Gly Leu Val Ala Thr Val 425 Tyr Ser Val Trp Leu Phe Tyr Ala Ala Glu Pro Gln Phe Val Leu Phe 440 Gly Ala Met Ala Met Leu Pro Gly Leu Ile Pro Tyr Val Trp Thr Arg 455 Ile Tyr Arg Gly Glu Gln Val Phe Asn Arg Phe Glu Ile Gly Val Val 470 475 Val Val Leu Val Val Ala Ala Ser Ala Gly Val Ile Gly Leu Val Asn 485 490 Gly Ser Leu Ser Leu 500 <210> 53 <211> 822 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(799) <223> RXA01394 <400> 53 gagcaaagtg tccagttgaa tggggttcat gaagctatat taaaccatgt taagaaccaa 60 tcattttact taagtacttc cataggtcac gatggtgatc atg gaa atc ttc att 115 Met Glu Ile Phe Ile 163 aca ggt ctg ctt ttg ggg gcc agt ctt tta ctg tcc atc gga ccg cag Thr Gly Leu Leu Gly Ala Ser Leu Leu Ser Ile Gly Pro Gln 15 aat gta ctg gtg att aaa caa gga att aag cgc gaa gga ctc att gcg 211 Asn Val Leu Val Ile Lys Gln Gly Ile Lys Arg Glu Gly Leu Ile Ala 259 gtt ctt ctc gtg tgt tta att tct gac gtc ttt ttg ttc atc gcc ggc Val Leu Leu Val Cys Leu Ile Ser Asp Val Phe Leu Phe Ile Ala Gly 45 acc ttg ggc gtt gat ctt ttg tcc aat gcc gcg ccg atc gtg ctc gat 307 Thr Leu Gly Val Asp Leu Leu Ser Asn Ala Ala Pro Ile Val Leu Asp 55 att atg cgc tgg ggt ggc atc gct tac ctg tta tgg ttt gcc gtc atg 355 Ile Met Arg Trp Gly Gly Ile Ala Tyr Leu Leu Trp Phe Ala Val Met 70 75 80

-	gcg Ala		-		_			-		_			_			403
-	gaa Glu		_					-	-	_		_	~ -	~ ~	_	451
-	gtg Val	-		_	_									-	_	499
-	aag Lys 135						-		_	_	_				-	547
	tgg Trp	_		_				-	-							595
	gtc Val						-							-	-	643
	gcg Ala			_	_	_				-	_					691
-	gca Ala	-	_		-	_	-		-		-			_		739
	aac Asn 215		-		_	_		_		-	_	_			-	787
_	ttg Leu	_		tagi	tttt	ege (	gggtt	ttg	ga at	cc						822
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Ser	Ile	Gly	Pro 20	Gln	Asn	Val	Leu	Val 25	Ile	Lys	Gln	Gly	Ile 30	Lys	Arg	
Glu	Gly	Leu 35	Ile	Ala	Val	Leu	Leu 40	Val	Cys	Leu	Ile	Ser 45	Asp	Val	Phe	
Leu	Phe 50	Ile	Ala	Gly	Thr	Leu 55	Gly	Val	Asp	Leu	Leu 60	Ser	Asn	Ala	Ala	
Pro	Ile	Val	Leu	Asp	Ile	Met	Arg	Trp	Gly	Gly	Ile	Ala	Tyr	Leu	Leu	

6	5				70					75					80	
Tr	p Phe	Ala	Val	Met 85	Ala	Ala	Lys	Asp	Ala 90	Met	Thr	Asn	Lys	Val 95	Glu	
Al	a Pro	Gln	Ile 100	Ile	Glu	Glu	Thr	Glu 105	Pro	Thr	Val	Pro	Asp 110	Asp	Thr	
Pr	o Leu	Gly 115	Gly	Ser	Ala	Val	Ala 120	Thr	Asp	Thr	Arg	Asn 125	Arg	Val	Arg	
Va	l Glu 130	Val	Ser	Val	Asp	Lys 135	Gln	Arg	Val	Trp	Val 140	Lys	Pro	Met	Leu	
Ме 14	t Ala 5	Ile	Val	Leu	Thr 150	Trp	Leu	Asn	Pro	Asn 155	Ala	Tyr	Leu	Asp	Ala 160	
Ph	e Val	Phe	Ile	Gly 165	Gly	Val	Gly	Ala	Gln 170	Tyr	Gly	Asp	Thr	Gly 175	Arg	
Tr	p Ile	Phe	Ala 180	Ala	Gly	Ala	Phe	Ala 185	Ala	Ser	Leu	Ile	Trp 190	Phe	Pro	
Le	u Val	Gly 195	Phe	Gly	Ala	Ala	Ala 200	Leu	Ser	Arg	Pro	Leu 205	Ser	Ser	Pro	
Ly	s Val 210	Trp	Arg	Trp	Ile	Asn 215	Val	Val	Val	Ala	Val 220	Val	Met	Thr	Ala	
Le ²	u Ala 5	Ile	Lys	Leu	Met 230	Leu	Met	Gly								
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tt	taacc	ccc a	aaat	gagg	ga aç	gaag	gtaad	c cti	tgaad	ctct		agc Ser				115
	a gct r Ala	_			_	_					_		_	_	-	163
_	t act l Thr			_	_			-								211
_	a gtc u Val		-		_	_	-	_					_			259

	~ ~			ggt Gly	-			_			-		-			307
_	_			gcc Ala												355
	_		_	gga Gly 90				-					-			403
-	-	-	-	tct Ser			-	-				-				451
				ccg Pro												499
	-	_	_	aca Thr		-			-			_				547
				cca Pro				-		_	-	_	_	_	_	595
		_		ttg Leu 170	-	-	-	-	_	_		-		_		643
-	_		_	atc Ile			_			-					-	691
•				ctt Leu	_			_	_							739
	-			cat His	-						_					787
-		-		ggc Gly					-							835
		_	_	gta Val 250	-	_			_	-			-	_	_	883
_		~	_	ctg Leu	-	_	_				_				-	931
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<212> PRT

<213> Corynebacterium glutamicum

<400> 56

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Val Gly Val Ala Met Val Thr Pro Phe Thr Glu Ser Gly Asp Ile Asp 20 25 30

Ile Ala Ala Gly Arg Glu Val Ala Ala Tyr Leu Val Asp Lys Gly Leu 35 40 45

Asp Ser Leu Val Leu Ala Gly Thr Thr Gly Glu Ser Pro Thr Thr Thr 50 55 60

Ala Ala Glu Lys Leu Glu Leu Leu Lys Ala Val Arg Glu Glu Val Gly 65 70 75 80

Asp Arg Ala Lys Leu Ile Ala Gly Val Gly Thr Asn Asn Thr Arg Thr 85 90 95

Ser Val Glu Leu Ala Glu Ala Ala Ala Ser Ala Gly Ala Asp Gly Leu 100 105 110

Leu Val Val Thr Pro Tyr Tyr Ser Lys Pro Ser Gln Glu Gly Leu Leu 115 120 125

Ala His Phe Gly Ala Ile Ala Ala Thr Glu Val Pro Ile Cys Leu 130 135 140

Tyr Asp Ile Pro Gly Arg Ser Gly Ile Pro Ile Glu Ser Asp Thr Met 145 150 155 160

Arg Arg Leu Ser Glu Leu Pro Thr Ile Leu Ala Val Lys Asp Ala Lys
165 170 175

Gly Asp Leu Val Ala Ala Thr Ser Leu Ile Lys Glu Thr Gly Leu Ala 180 185 190

Trp Tyr Ser Gly Asp Asp Pro Leu Asn Leu Val Trp Leu Ala Leu Gly 195 200 205

Gly Ser Gly Phe Ile Ser Val Ile Gly His Ala Ala Pro Thr Ala Leu 210 215 220

Arg Glu Leu Tyr Thr Ser Phe Glu Glu Gly Asp Leu Val Arg Ala Arg 225 230 235 240

Glu Ile Asn Ala Lys Leu Ser Pro Leu Val Ala Ala Gln Gly Arg Leu 245 250 255

Gly Gly Val Ser Leu Ala Lys Ala Ala Leu Arg Leu Gln Gly Ile Asn

265

260

270

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cac gtc tac tgt gtg gac cgt ctt ccc cgc atg gtc gac tat gtg gtt
                                                                   595
His Val Tyr Cys Val Asp Arg Leu Pro Arg Met Val Asp Tyr Val Val
                    155
                                                                   643
ccc cct gga gtc cgc atc tcc gaa gca gaa cgc gtg cgc cta ggt gca
Pro Pro Gly Val Arg Ile Ser Glu Ala Glu Arg Val Arg Leu Gly Ala
                170
                                    175
                                                                   691
tac ctt gct ccg ggt acc tct gtg ctg cgt gaa ggt ttc gtg tct ttc
Tyr Leu Ala Pro Gly Thr Ser Val Leu Arg Glu Gly Phe Val Ser Phe
                                190
            185
                                                                   739
aac too ggc acc ttg ggt gcc gca aag gtg gaa ggc cgc ctg agt tcc
Asn Ser Gly Thr Leu Gly Ala Ala Lys Val Glu Gly Arg Leu Ser Ser
                            205
                                                                   787
qqt qtq qtc atc ggt gaa ggt tcc gag att gga ctg tct tct act att
Gly Val Val Ile Gly Glu Gly Ser Glu Ile Gly Leu Ser Ser Thr Ile
                        220
                                                                   835
cag tee eeg aga gat gaa cag ege ege egt tig eeg tig age ate gge
Gln Ser Pro Arg Asp Glu Gln Arg Arg Leu Pro Leu Ser Ile Gly
                    235
                                        240
caa aac tgc aac ttt ggt gtc agc tcc gga atc atc gga gtc agt ctg
                                                                   883
Gln Asn Cys Asn Phe Gly Val Ser Ser Gly Ile Ile Gly Val Ser Leu
                                    255
                                                                   931
gga gac aat tgc gac atc gga aat aac att gtc ttg gat gga gat acc
Gly Asp Asn Cys Asp Ile Gly Asn Asn Ile Val Leu Asp Gly Asp Thr
            265
                                270
                                                                   979
ccc att tgg ttc gca gcc gat gag gag tta cgc act atc gac tcc atc
Pro Ile Trp Phe Ala Ala Asp Glu Glu Leu Arg Thr Ile Asp Ser Ile
                            285
        280
gaa ggc caa gca aat tgg tca atç aag cgt gaa tcc ggc ttc cat gag
Glu Gly Gln Ala Asn Trp Ser Ile Lys Arg Glu Ser Gly Phe His Glu
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<211> 316

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<400> 58

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Ala Met Asp Gly Thr Ile Leu Asp Thr Trp Tyr Pro Glu Pro Gln Ile 20 25 30

Phe Asn Pro Asp Gln Trp Ala Glu Arg Tyr Pro Leu Glu Val Gly Thr

35 40 45

Thr Arg Leu Gly Ala Asn Glu Leu Thr Pro Arg Met Leu Gln Leu Val
50 60

Lys Leu Asp Gln Asp Arg Leu Val Glu Gln Val Ala Val Arg Thr Val 65 70 75 80

Ile Pro Asp Leu Ser Gln Pro Pro Val Asp Ala His Asp Val Tyr Leu
85 90 95

Arg Leu His Leu Leu Ser His Arg Leu Val Arg Pro His Glu Met His  $100 \hspace{1cm} 105 \hspace{1cm} 110$ 

Met Gln Asn Thr Leu Glu Leu Leu Ser Asp Val Val Trp Thr Asn Lys 115 120 125

Gly Pro Cys Leu Pro Glu Asn Phe Glu Trp Val Arg Gly Ala Leu Arg 130 140

Ser Arg Gly Leu Ile His Val Tyr Cys Val Asp Arg Leu Pro Arg Met 145 150 155 160

Val Asp Tyr Val Val Pro Pro Gly Val Arg Ile Ser Glu Ala Glu Arg 165 170 175

Val Arg Leu Gly Ala Tyr Leu Ala Pro Gly Thr Ser Val Leu Arg Glu 180 185 190

Gly Phe Val Ser Phe Asn Ser Gly Thr Leu Gly Ala Ala Lys Val Glu 195 200 205

Gly Arg Leu Ser Ser Gly Val Val Ile Gly Glu Gly Ser Glu Ile Gly 210 215 220

Leu Ser Ser Thr Ile Gln Ser Pro Arg Asp Glu Gln Arg Arg Arg Leu 225 230 235 240

Pro Leu Ser Ile Gly Gln Asn Cys Asn Phe Gly Val Ser Ser Gly Ile 245 250 255

Ile Gly Val Ser Leu Gly Asp Asn Cys Asp Ile Gly Asn Asn Ile Val  $260 \hspace{1cm} 265 \hspace{1cm} 270 \hspace{1cm}$ 

Leu Asp Gly Asp Thr Pro Ile Trp Phe Ala Ala Asp Glu Glu Leu Arg 275 280 285

Thr Ile Asp Ser Ile Glu Gly Gln Ala Asn Trp Ser Ile Lys Arg Glu 290 295 300

Ser Gly Phe His Glu Pro Val Ala Arg Leu Lys Ala 305 310 315

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<211> 1296

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<220>

<221> CDS

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tgc gat gag tac ggc atc ttg atg atc acc gat gaa gtc cag act ggc
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Cys Asp Glu Tyr Gly Ile Leu Met Ile Thr Asp Glu Val Gln Thr Gly
    215
                        220
gtt ggc cgt acc ggc gat ttc ttt gca cat cag cac gat ggc gtt gtt
                                                                   835
Val Gly Arg Thr Gly Asp Phe Phe Ala His Gln His Asp Gly Val Val
230
                    235
                                        240
ccc gat gtg gtg acc atg gcc aag gga ctt ggc ggc ggt ctt ccc atc
                                                                   883
Pro Asp Val Val Thr Met Ala Lys Gly Leu Gly Gly Gly Leu Pro Ile
                250
                                    255
ggt gct tgt ttg gcc act ggc cgt gca gct gaa ttg atg acc cca ggc
Gly Ala Cys Leu Ala Thr Gly Arg Ala Ala Glu Leu Met Thr Pro Gly
            265
                                270
aag cac ggc acc act ttc ggt ggc aac cca gtt gct tgt gca gct gcc
Lys His Gly Thr Thr Phe Gly Gly Asn Pro Val Ala Cys Ala Ala Ala
        280
                            285
aag gca gtg ctg tet gtt gtc gat gac gct ttc tgc gca gaa gtt gcc
Lys Ala Val Leu Ser Val Val Asp Asp Ala Phe Cys Ala Glu Val Ala
cgc aag ggc gag ctg ttc aag gaa ctt ctt gcc aag gtt gac ggc gtt
Arg Lys Gly Glu Leu Phe Lys Glu Leu Leu Ala Lys Val Asp Gly Val
gta gac gtc cgt ggc agg ggc ttg atg ttg ggc gtg gtg ctg gag cgc
1123
Val Asp Val Arg Gly Arg Gly Leu Met Leu Gly Val Val Leu Glu Arg
gac gtc gca aag caa gct gtt ctt gat ggt ttt aag cac ggc gtt att
1171
Asp Val Ala Lys Gln Ala Val Leu Asp Gly Phe Lys His Gly Val Ile
            345
                                350
ttg aat gca ccg gcg gac aac att atc cgt ttg acc ccg ccg ctg gtg
1219
Leu Asn Ala Pro Ala Asp Asn Ile Ile Arg Leu Thr Pro Pro Leu Val
                            365
atc acc gac gaa gaa atc gca gac gca gtc aag gct att gcc gag aca
Ile Thr Asp Glu Glu Ile Ala Asp Ala Val Lys Ala Ile Ala Glu Thr
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atc gca taaaggactc aaacttatga ctt
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Ile Ala
390
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<212> PRT
<213> Corynebacterium glutamicum
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Lys Val Asp Gly Val Val Asp Val Arg Gly Arg Gly Leu Met Leu Gly 325 330 Val Val Leu Glu Arg Asp Val Ala Lys Gln Ala Val Leu Asp Gly Phe Lys His Gly Val Ile Leu Asn Ala Pro Ala Asp Asn Ile Ile Arg Leu 360 Thr Pro Pro Leu Val Ile Thr Asp Glu Glu Ile Ala Asp Ala Val Lys 375 Ala Ile Ala Glu Thr Ile Ala <210> 61 <211> 1008 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(985) <223> RXC00733 <400> 61 acggcgaggt tgtcggtatt ggaacgcaca cgaatttgct gaacacgtgc ggtacctacc 60 gtgaaattgt tgaatcccaa gagactgcgc aggcgcaatc atg agt aat act gca Met Ser Asn Thr Ala 163 ggc ccc cgc ggg cgt tcc cat cag gca gac gcc gcg ccg aat caa aag Gly Pro Arg Gly Arg Ser His Gln Ala Asp Ala Ala Pro Asn Gln Lys gca cag aat ttc gga cca tct gcc aaa agg ctt ttc gga att cta ggc 211 Ala Gln Asn Phe Gly Pro Ser Ala Lys Arg Leu Phe Gly Ile Leu Gly cat gac cgt aac acc tta att ttt gtt atc ttc cta gcc gtc ctg agc 259 His Asp Arg Asn Thr Leu Ile Phe Val Ile Phe Leu Ala Val Leu Ser 45 307 gtt gga ctt acc gtc ttg ggc cca tgg ttg ctg ggt aaa gcc acc aac Val Gly Leu Thr Val Leu Gly Pro Trp Leu Leu Gly Lys Ala Thr Asn 60 gtg gtg ttt gaa gga ttc cta tct aag cgc atg ccg gcl ggl gcg tca 355 Val Val Phe Glu Gly Phe Leu Ser Lys Arg Met Pro Ala Gly Ala Ser 80 403 aaq qaa qat atc atc qcq caq ttq caq gct gca ggt aaa cat aat cag Lys Glu Asp Ile Ile Ala Gln Leu Gln Ala Ala Gly Lys His Asn Gln 95 gct tcc atg atg gaa gac atg aac ctt gtt cca ggc tca ggc att gat Ala Ser Met Met Glu Asp Met Asn Leu Val Pro Gly Ser Gly Ile Asp 105 110

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Phe Glu Lys Leu Ala Met Ile Leu Gly Leu Val Ile Gly Ala Tyr Leu
        120
                             125
atc ggt agc ctg ttg ttg ttc cag gcg cgg atg ctc aac cgc atc
                                                                    547
Ile Gly Ser Leu Leu Ser Leu Phe Gln Ala Arg Met Leu Asn Arg Ile
    135
                         140
                                             145
gtg caa agt gcc atg cac cgg ctg cgc atg gag gtg gag gaa aaa atc
                                                                    595
Val Gln Ser Ala Met His Arg Leu Arg Met Glu Val Glu Glu Lys Ile
                    155
cac cgc cta ccg ctg agc tat ttc gat tcc atc aaa cgt ggt gat ctg
                                                                    643
His Arg Leu Pro Leu Ser Tyr Phe Asp Ser Ile Lys Arg Gly Asp Leu
                170
                                     175
ctt agc cgt gtg acc aac gat gtg gat aat atc ggt caa tcc ctg caa
                                                                    691
Leu Ser Arg Val Thr Asn Asp Val Asp Asn Ile Gly Gln Ser Leu Gln
                                 190
caa acc ttg tca cag gcg atc act tcc cta ctg acc gtc atc ggt gtg
                                                                    739
Gln Thr Leu Ser Gln Ala Ile Thr Ser Leu Leu Thr Val Ile Gly Val
                             205
ttg gtg atg atg ttt atc atc tcc cca ctg ctc gca ctc gtg gcg ctg
                                                                    787
Leu Val Met Met Phe Ile Ile Ser Pro Leu Leu Ala Leu Val Ala Leu
gta too att cog gto acc atc gtg gtc act gtg gtg gtt gcg agc cgt
                                                                    835
Val Ser Ile Pro Val Thr Ile Val Val Thr Val Val Val Ala Ser Arg
                    235
                                         240
tcc cag aaa ctc ttt gcg gaa cag tgg aag cag acc ggt att ttg aat
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Ser Gln Lys Leu Phe Ala Glu Gln Trp Lys Gln Thr Gly Ile Leu Asn
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                                     255
gcg cgc ctg gag gaa acc tac tct ggc cac gcc gtg gtt aag gtt ttc
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Ala Arg Leu Glu Glu Thr Tyr Ser Gly His Ala Val Val Lys Val Phe
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                                 270
<u>qqa cac caa aaq qat qtt caa qaa qca ttc qaq qaa qaa aat caa qct</u>
Gly His Gln Lys Asp Val Gln Glu Ala Phe Glu Glu Glu Asn Gln Ala
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1008
Cys Val
    295
<210> 62
<211> 295
<212> PRT
<213> Corynebacterium glutamicum
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Ala Pro Asn Gln Lys Ala Gln Asn Phe Gly Pro Ser Ala Lys Arg Leu
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20 25 30

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Leu Ala Val Leu Ser Val Gly Leu Thr Val Leu Gly Pro Trp Leu Leu 50 60

Gly Lys Ala Thr Asn Val Val Phe Glu Gly Phe Leu Ser Lys Arg Met
65 70 75 80

Pro Ala Gly Ala Ser Lys Glu Asp Ile Ile Ala Gln Leu Gln Ala Ala 85 90 95

Gly Ser Gly Ile Asp Phe Glu Lys Leu Ala Met Ile Leu Gly Leu Val 115 120 125

Ile Gly Ala Tyr Leu Ile Gly Ser Leu Leu Ser Leu Phe Gln Ala Arg 130 135 140

Met Leu Asn Arg Ile Val Gl<br/>n Ser Ala Met His Arg Leu Arg Met Glu 145  $\,$  150  $\,$  155  $\,$  160

Val Glu Glu Lys Ile His Arg Leu Pro Leu Ser Tyr Phe Asp Ser Ile 165 170 175

Lys Arg Gly Asp Leu Leu Ser Arg Val Thr Asn Asp Val Asp Asn Ile 180 185 190

Gly Gln Ser Leu Gln Gln Thr Leu Ser Gln Ala Ile Thr Ser Leu Leu 195 200 205

Thr Val Ile Gly Val Leu Val Met Met Phe Ile Ile Ser Pro Leu Leu 210 215 220

Ala Leu Val Ala Leu Val Ser Ile Pro Val Thr Ile Val Val Thr Val 225 230 235 240

Val Val Ala Ser Arg Ser Gln Lys Leu Phe Ala Glu Gln Trp Lys Gln 245 250 255

Thr Gly Ile Leu Asn Ala Arg Leu Glu Glu Thr Tyr Ser Gly His Ala 260 265 270

Val Val Lys Val Phe Gly His Gln Lys Asp Val Gln Glu Ala Phe Glu 275 280 285

Glu Glu Asn Gln Ala Cys Val 290 295

<210> 63

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<212> DNA

<213> Corynebacterium glutamicum

<220>

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<213> Corynebacterium glutamicum

<400> 64

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Thr Val Arg Asp Gly Asp Leu Ile Ile Leu Ser Ser Ser Leu Val Pro 35 40 45

Gly Asn Glu Glu Ala Val Phe Gly Val Ile Asn Met Leu Ala Gln Ile 50 60

Gly Ala Thr Val Val Thr Gly Arg Asp Ala Lys Val His Thr Ser Gly 65 70 75 80

His Gly Tyr Ser Gly Glu Leu Leu Phe Leu Tyr Asn Ala Ala Arg Pro 90 Lys Asn Ala Met Pro Val His Gly Glu Trp Arg His Leu Arg Ala Asn 105 Lys Glu Leu Ala Ile Ser Thr Gly Val Asn Arg Asp Asn Val Val Leu 120 115 Ala Gln Asn Gly Val Val Val Asp Met Val Asn Gly Arg Ala 135 <210> 65 <211> 1066 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1066) <223> RXC00866 <400> 65 gcatcaacgt aggagatect cgacttecaa ttatggetee aaatgageag gaacttgagg 60 ctctccgaga agacatgaaa aaagctggag ttctataaat atg aat gat tcc cga Met Asn Asp Ser Arg 1 aat cgc ggc cgg aag gtt acc cgc aag gcg ggc cca cca gaa gct ggt Asn Arg Gly Arg Lys Val Thr Arg Lys Ala Gly Pro Pro Glu Ala Gly 15 10 cag gaa aac cat ctg gat acc cct gtc ttt cag gca cca gat gct tcc 211 Gln Glu Asn His Leu Asp Thr Pro Val Phe Gln Ala Pro Asp Ala Ser 30 25 tct aac cag agc gct gta aaa gct gag acc gcc gga aac gac aat cgg 259 Ser Asn Gln Ser Ala Val Lys Ala Glu Thr Ala Gly Asn Asp Asn Arg 45 40 gat gct gcg caa ggt gct caa gga tcc caa gat tct cag ggt tcc cag 307 Asp Ala Ala Gln Gly Ala Gln Gly Ser Gln Asp Ser Gln Gly Ser Gln 60 55 aac gct caa ggt tcc cag aac cgc gag tcc gga aac aac aac cgc aac Asn Ala Gln Gly Ser Gln Asn Arg Glu Ser Gly Asn Asn Asn Arg Asn 70 75 403 cgt tcc aac aac cgt cgc ggt ggt cgt gga cgt cgt gga tcc gga Arg Ser Asn Asn Arg Arg Gly Gly Arg Gly Arg Gly Ser Gly 90 aac gcc aat gag ggc gcg aac aac agc ggt aac cag aac cgt cag Asn Ala Asn Glu Gly Ala Asn Asn Ser Gly Asn Gln Asn Arg Gln 105 115 ggc gga aac cgt ggc aac cgc ggt ggc gga cgc cga aac gtt gtt aag Gly Gly Asn Arg Gly Asn Arg Gly Gly Gly Arg Arg Asn Val Lys

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atc ggt Ile Gly	_		_												643
gtg gac Val Asp															691
ctg att Leu Ile			-												739
gat gca Asp Ala 215	_		_					_	_				-		787
ccc tgg Pro Trp 230	_	_	_	_	-		_				_	-			835
ttc acc Phe Thr		-													883
ccg aag Pro Lys	_		-	_			_				_	_		_	931
ttc aac Phe Asn		_			_	_									979
ggt ctt 1027	get	acc	aag	act	cct	gct	ggt	ttg	gtc	atc	cac	acc	ggt	gac	
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gtg a																691
ttg (																739
ttt q Phe i																787
cat ( His 1 230																835
cag ( Gln (																883
ttt ( Phe																931
atg Met																979
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Ile		Glu	Val 345	Ile	Asp	Leu	Gln	Pro 350	Glu	Arg	Thr	Asp	Pro 355	Ala	His	
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Phe Asp Ser Asn Gly His Arg Thr Arg Phe Asp Asp Leu Thr His Ser 425 430 435

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Arg Pro Trp Leu Thr Ser Phe Thr Val Ile Ser Ala Leu Ala Ala Thr  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

bed file Gld bed file bed file bed bed file Gly Gly Ala lie Asp lie 50 55 60

Ala Leu Gly Asn Thr Gly Asp Thr Leu Thr Thr Asp Leu Leu Asp Arg 65 70 75 80

Phe Thr Pro Ser Gly Leu Ser Val Leu Thr Ser Val Ile Ala Leu Ile 85 90 95

Val Leu Leu Ala Leu Leu Arg Tyr Ala Ser Gln Phe Gly Arg Tyr 100 105 110

Thr Ala Gly Lys Leu Ser Met Gly Val Gln His Asp Val Arg Leu Lys
115 120 125

Thr Met Arg Ser Leu Gln Asn Leu Asp Gly Pro Gly Gln Asp Ser Ile 130 135 140

Arg Thr Gly Gln Val Val Ser Arg Ser Ile Ser Asp Ile Asn Met Val

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Ile	Ile	Ala 195	Ala	Val	Leu	Val	Pro 200	Leu	Leu	Leu	Trp	Ala 205	Val	Ala	Tyr
Ser	Arg 210	Lys	Ala	Leu	Phe	Ala 215	Ser	Thr	Trp	Ser	Ala 220	Gln	Gln	Lys	Ala
Ala 225	Asp	Leu	Thr	Thr	His 230	Val	Glu	Glu	Thr	Val 235	Thr	Gly	Ile	Arg	Val 240
Val	Lys	Ala	Phe	Ala 245	Gln	Glu	Asp	Arg	Glu 250	Thr	Asp	Lys	Leu	Asp 255	Leu
Thr	Ala	Arg	Glu 260	Leu	Phe	Ala	Gln	Arg 265	Met	Arg	Thr	Ala	Arg 270	Leu	Thr
Ala	Lys	Phe 275	Ile	Pro	Met	Val	Glu 280	Gln	Leu	Pro	Gln	Leu 285	Ala	Leu	Val
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Val 305	Gly	Thr	Phe	Val	Ala 310	Phe	Ser	Ser	Tyr	Leu 315	Thr	Ser	Leu	Ser	Ala 320
Val	Ala	Arg	Ser	Leu 325	Ser	Gly	Met	Leu	Met 330	Arg	Val	Gln	Leu	Ala 335	Leu
Ser	Ser	Val	Glu 340	Arg	Ile	Phe	Glu	Val 345	Ile	Asp	Leu	Gln	Pro 350	Glu	Arg
Thr	Asp	Pro 355	Ala	His	Pro	Leu	Ser 360	Leu	Pro	Asp	Thr	Pro 365	Leu	Gly	Leu
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Gly	Lys	Thr	Met	Ala 405	Val	Gln	Leu	Ala	Gly 410	Asn	Phe	Tyr	Gln	Pro 415	Asp
Ser	Gly	His	Ile 420	Ala	Phe	Asp	Ser	Asn 425	Gly	His	Arg	Thr	Arg 430	Phe	Asp
Asp	Leu	Thr 435	His	Ser	Asp	Ile	Arg 440	Arg	Asn	Leu	Ile	Ala 445	Val	Phe	Asp
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Asp Pro Tyr Arg Met Val Gln Gln Leu Arg Arg Lys Leu Ser Arg Phe
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Val Glu Gln Lys Trp Lys Arg Gln Pro Val Ile Met Pro Thr Val Ile
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ccg atg act gcg gaa acc acg cac atc ggt gac gat gag gtt cgc gct
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agc tgg Ser Trp 230	_	_			_		_		_	-					835
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tgc acc Cys Thr	_			_			-	~		_	-		_		979
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gtt gtg 1267	atg	gct	tct	gaa	tcg	gga	gtg	ttg	gac	ttg	agg	gag	gag	agc	
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Val Val 390	Lys	Arg	Thr	Arg 395	Val	Gln	Pro	Gly	Arg 400	Met	Phe	Leu	Val	Asp 405	
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Lys Thr Gly Arg Asp Val Val Ile Ala Ala Leu Leu Gly Ala Glu Glu 1095 1100 1105

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Pro Asp Leu Arg Ser Lys Phe Thr Gly Lys Ala Glu His Val Val Asn 1145 1150 1155

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Val Arg Cys Thr Lys Thr Gln Glu His Ser Leu Glu Lys Ala Leu Asp 1225 1230 1235

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Ile	Lys	Gln	Lys 420	Leu	Ser	Glu	Ala	Gln 425	Pro	Tyr	Gly	Glu	Trp 430	Ile	Arg
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Glu	Ala	Ile	Gly	Ser 485	Met	Gly	Ser	Asp	Thr 490	Pro	Ile	Ala	Ala	Leu 495	Ser
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Ala	Ile 610	Arg	Asn	Gly	Lys	Thr 615	Leu	Ile	Val	Leu	Ser 620	Asp	Arg	Glu	Ser
Asp 625	Glu	Arg	Met	Ala	Pro 630	Ile	Pro	Ala	Leu	Leu 635	Leu	Thr	Ser	Ala	Val 640
His	Gln	Tyr	Leu	Val 645	Gln	Gln	Arg	Thr	Arg 650	Thr	Gln	Cys	Ser	Leu 655	Val
Val	Glu	Ser	Gly 660	Asp	Ala	Arg	Glu	Val 665	His	His	Leu	Ala	Met 670	Leu	Ile
Gly	Phe	Gly 675	Ala	Asp	Ala	Ile	Asn 680	Pro	Tyr	Met	Ala	Phe 685	Glu	Thr	Ile
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- Glu His Val Val Asn Phe Phe Thr Phe Ile Ala Gln Glu Val Arg Glu
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- Phe Pro Thr Gln Asp Val Arg Cys Thr Lys Thr Gln Glu His Ser Leu 1220 1225 1230
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Val	Asp	Val	Val	Pro 144		Ser	Ala	Asp	Asp	Leu 50	Thr	Trp	Ala	Asp (	Glu 55	
Leu	Ile	Ala	Arg 146		Arg	Glu	Leu	Thr 14	Gly 65	Ser	Glu	Thr	Lys 1	Leu . 470	Arg	
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							atc Ile							691
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tcg Ser	-		-				tcg ser	 					 	787
	215					220				225				
							tac Tyr							835
				_			gaa Glu		_	_	_	-		883
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Ala Gly Ala Glu Lys Asn Thr Gly Asp Gly Ala Gly Ile Leu Met Gln 50 60

Ile Pro Asp Gly Phe Tyr Arg Glu Val Ser Gly Ile Glu Leu Pro Glu65707580

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Pro Leu Val Val Glu Gly Cys Ile Met Met Arg Val Cys His Leu Asp 345 Thr Cys Pro Val Gly Ile Ala Thr Gln Asn Pro Asp Leu Arg Ser Lys 360 Phe Thr Gly Lys Ala Glu His Val Val Asn Phe Phe Thr Phe Ile Ala 375 Gln Glu Val Arg Glu Tyr Leu Ala Gln Leu Gly Phe Arg Ser Ile Asp 390 395 Glu Ala Val Gly Gln Ala Gln Val Leu Arg Lys Arg Ser Gly Ile Pro 410 Ala Asp Ser Arg Ala Ala His Leu Asp Leu Ser Pro Ile Phe Ile 420 425 <210> 77 <211> 866 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (1)..(843) <223> FRXA00367 <400> 77 cac age eta gaa aaa gee etg gae aac gea tit att gat aag get teg His Ser Leu Glu Lys Ala Leu Asp Asn Ala Phe Ile Asp Lys Ala Ser 5 gac acg atc acc cgt gcc gca gcg ggt gtg gaa acc agc att gtt att 96 Asp Thr Ile Thr Arg Ala Ala Gly Val Glu Thr Ser Ile Val Ile 20 25 gat agc tcc atc agc aac gtc aac cgt tca gtt ggc acg atg ctg ggt 144 Asp Ser Ser Ile Ser Asn Val Asn Arg Ser Val Gly Thr Met Leu Gly 35 tet gea gte age ege gtg get ggt gee eaa ggt ttg eea gae gge ace Ser Ala Val Ser Arg Val Ala Gly Ala Gln Gly Leu Pro Asp Gly Thr 5.0 ate ace ttg aat ett caa gge tge gee ggt aac tee ttt gge geg tte 240 Ile Thr Leu Asn Leu Gln Gly Cys Ala Gly Asn Ser Phe Gly Ala Phe 65 atc cca cga ggc atc acc atc acc ctc acc ggc gat gcc aat gac ttt 288 Ile Pro Arg Gly Ile Thr Ile Asn Leu Thr Gly Asp Ala Asn Asp Phe 85 gtg ggc aag gga tta tct ggc gga aag att gtg atc aag cct tcc gct 336 Val Gly Lys Gly Leu Ser Gly Gly Lys Ile Val Ile Lys Pro Ser Ala cag gct ccg aag cag ctg aag aac aat cca aat atc att gcc gga aac 384 Gln Ala Pro Lys Gln Leu Lys Asn Asn Pro Asn Ile Ile Ala Gly Asn 120

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	gtt Val															595
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	gcc Ala															835
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aag gac Lys Asp															979
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His Asn Val Leu Asp Lys Pro Arg Leu Arg Leu Leu Gly Asn Ile Glu 65 70 75 80

Ile Gly Lys Asp Ile Thr Val Glu Glu Leu Arg Asp Tyr Tyr Asp Ala 85 90 95

Val Val Phe Ser Thr Gly Ala Val Ala Asp Arg Asp Leu Asn Ile Pro  $100 \,\,$ 

Gly Ile Glu Ala Glu Gly Ser Phe Gly Ala Gly Glu Phe Val Gly Phe
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Ser Val Ala Val Ile Gly Val Gly Asn Val Gly Leu Asp Val Ala Arg 145 150 155 160

Ile Leu Ala Lys Thr Gly Asp Glu Leu Lys Val Thr Glu Ile Ser Asp 165 170 175

Asn Val Tyr Asp Ser Leu Lys Glu Asn Lys Ala Thr Glu Val His Val 180 185 190

Phe Gly Arg Arg Gly Pro Ala Gln Val Lys Phe Thr Pro Gln Glu Leu 195 200 205

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Asp Ile Asp Tyr Asp Gly Ala Ser Glu Glu Ala Arg Arg Ala Ser Lys 225 230 235 240

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Pro Lys Asp Ala Pro His Thr Leu Gln Ile His Leu Phe Glu Asn Pro

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gtc Val	gto Val	ggt LGly 4(	t too y Ser )	ggc Gly	ccc Pro	gct Ala	ggc Gly 45	Leu	gcc Ala	gcc Ala	gcg Ala	cag Gln 50	cag Gln	ctc Leu	acc Thr	259
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Asp Asp Arg Leu Gly Gly Leu Met Arg Tyr Gly Val Pro Glu Tyr Lys
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Met Glu Asn Arg Trp Ile Asp Arg Ile Glu Gln Met Glu Ala Glu 85 90 95

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136

105

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100

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Asp Arg Leu His Ala Thr Asn Asn Phe Pro Glu Phe Thr Gly Arg Leu 85 90 95

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931

979

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Gly Leu Ile Gln Arg Leu Cys Glu Pro Glu Arg Gln Leu Ile Phe Arg 50 55 60

Val Pro Trp Val Asp Asp Gln Gly Gln Val His Val Asn Arg Gly Phe 65 70 75 80

Arg Val Gln Phe Asn Ser Ala Leu Gly Pro Tyr Lys Gly Gly Leu Arg 85 90 95

Phe His Pro Ser Val Asn Leu Gly Ile Val Lys Phe Leu Gly Phe Glu 100 105 110

Gln Ile Phe Lys Asn Ser Leu Thr Gly Leu Pro Ile Gly Gly Lys 115 120 125

Gly Gly Ser Asp Phe Asp Pro Lys Gly Lys Ser Asp Leu Glu Ile Met

Arg Phe Cys Gln Ser Phe Met Thr Glu Leu His Arg His Ile Gly Glu 145 150 155 160

Tyr Arg Asp Val Pro Ala Gly Asn Ile Gly Val Gly Gly His Glu Ile 165 170 175

Gly Tyr Leu Phe Gly His Tyr Arg Arg Met Ala Asn Gln His Glu Ser 180 185 190

Gly Val Leu Thr Gly Lys Gly Leu Thr Trp Gly Gly Ser Leu Val Arg 195 200 205

Thr Glu Ala Thr Gly Tyr Gly Cys Val Tyr Phe Val Ser Glu Met Ile 210 215 220

Lys Ala Lys Gly Glu Ser Ile Ser Gly Gln Lys Ile Ile Val Ser Gly 225 230 235 240

Ser Gly Asn Val Ala Thr Tyr Ala Ile Glu Lys Ala Gln Glu Leu Gly 245 250 255

Ala Thr Val Ile Gly Phe Ser Asp Ser Ser Gly Trp Val His Thr Pro 260 265 270

Asn Gly Val Asp Val Ala Lys Leu Arg Glu Ile Lys Glu Val Arg Arg 280 Ala Arg Val Ser Val Tyr Ala Asp Glu Val Glu Gly Ala Thr Tyr His 290 295 300 Thr Asp Gly Ser Ile Trp Asp Leu Lys Cys Asp Ile Ala Leu Pro Cys 310 Ala Thr Gln Asn Glu Leu Asn Gly Glu Asn Ala Lys Thr Leu Ala Asp Asn Gly Cys Arg Phe Val Ala Glu Gly Ala Asn Met Pro Ser Thr Pro Glu Ala Val Glu Val Phe Arg Glu Arg Asp Ile Arg Phe Gly Pro Gly Lys Ala Asa Asa Gly Gly Val Ala Thr Ser Ala Leu Glu Met Gln Gln Asn Ala Ser Arg Asp Ser Trp Ser Phe Glu Tyr Thr Asp Glu Arg 390 395 Leu Gln Val Ile Met Lys Asn Ile Phe Lys Thr Cys Ala Glu Thr Ala 410 Ala Glu Tyr Gly His Glu Asn Asp Tyr Val Val Gly Ala Asn Ile Ala Gly Phe Lys Lys Val Ala Asp Ala Met Leu Ala Gln Gly Val Ile <210> 95 <211> 1461 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101) .. (1438) <223> RAH00323 <400> 95 cgcgacatcc ttgagtaact ctgagaaaaa ctacccccga tgggagtata aaagtggcaa 60 atgcgcagtc gatgtcccat cgctgcgtag attagttttc atg aac agc gaa cag Met Asn Ser Glu Gln gaa ttt gta ctc agc gcc att gaa gaa cgc gac att aag ttt gtg cgt 163 Glu Phe Val Leu Ser Ala Ile Glu Glu Arg Asp Ile Lys Phe Val Arg 10 15 cta tgg ttc act gac att ctt ggc cac ttg aag tca gtg gtt gtg gct 211 Leu Trp Phe Thr Asp Ile Leu Gly His Leu Lys Ser Val Val Val Ala 25 30 cct gca gaa cta gag tct gcg ttg gaa gaa ggc atc gga ttc gat ggc Pro Ala Glu Leu Glu Ser Ala Leu Glu Glu Gly Ile Gly Phe Asp Gly

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cgc Arg 70	cca Pro	gat Asp	cca Pro	tcg Ser	aca Thr 75	ttc Phe	cag Gln	gtc Val	ctc Leu	cca Pro 80	cta Leu	gaa Glu	gcg Ala	ggc Gly	atc Ile 85	355
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cct Pro	gtc Val	gag Glu	ttc Phe 185	Ser	cac His	cat His	gaa Glu	act Thr 190	gca Ala	cct Pro	ggc Gly	cag Gln	caa Gln 195	gaa Glu	atc Ile	691
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tca Ser 230	Phe	atg Met	ccc Pro	: aag > Lys	cca Pro 235	Phe	caa Gln	gaa Glu	. cat . His	gca 3 Ala 240	ı Gly	tcc Ser	gcc Ala	atg Met	cac His 245	835
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gac Asp	gat Asp	tct Sei	t tac r Tyr 265	. Met	ctg Lev	g tco i Sei	aaa Lys	a acc Thr 270	Ala	a aaa a Lys	a caç s Glr	j tto n Phe	ato 116 275	e Ala	gga Gly	931
ato Ile	ttg E Lei	g cat i Hi: 280	s His	c gct s Ala	cca Pro	a gaa o Gli	a tto 1 Phe 285	e Thi	gc! Ala	t gte a Val	g aco l Thi	aac Asr 290	ı Glr	g tgg n Trp	g gtc Val	979

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cgc Arg 230	Phe	aac Asn	acc Thr	atg Met	ctc Leu 235	His	gcg Ala	gca Ala	gat Asp	gat Asp 240	Ile	cag Gln	acc Thr	ttc Phe	aag Lys 245	835
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caç Glr	g tco n Ser	cto Leu 280	rr	g aag Lys	gac Asp	ggc Gly	: aag / Lys 285	Pro	cto Leu	tto Phe	cac His	gat Asp 290	Glu	tcc Ser	ggc	979
		a ggc	ctg	g too	gad	ato	gcc	c gc	tac	: tac	ato	ggc	ggc	ato	ctg	
	295	5				300	)				305	)			. Leu	
10	75														tcc	
Hi:	s His	s Ala	a Gly	y Ala	a Val		ı Ala	a Phe	e Thi	r Asr 320		a Thi	. Le	ı Asr	Ser 325	

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ato		ago	c aac	c tog	g tto	: ttt	gag	g gto	g gat	gca	a gca	a ctt	cg(	c cca	a gaa	
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Leu	Leu 50	Trp	Gly	Leu	Ser	Gly 55	Ala	Gly	Asp	Pro	Asp 60	Val	Ala	Leu	Asn
Lev 65	Leu	Ile	Arg	Leu	Tyr 70	Gln	Ala	Leu	Glu	Ala 75	Ile	Gly	Glu	Asp	Ala 80
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Arg	Leu	Phe	Ala 100	Leu	Leu	Gly	Gly	Ser 105	Ser	Ala	Val	Gly	Asp 110	His	Leu
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Ala 145	Val	Leu	Glu	Val	Glu 150	Asp	Phe	Ser	Asp	Ala 155	His	Asn	Ile	Ala	Arg 160
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Ala	Glu 290	Leu	Ile	Arg	Ile	Gly 295	Ser	Asn	Ser	Phe	Phe 300	Glu	Val	Asp	Ala
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Ser	His	Met	Ala	Tyr 325	Tyr	Lys	Arg	Trp	Ala 330	Glu	Thr	Trp	Glu	Phe	Gln

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_			_					-		_		_	_	ctc Leu		384
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- Val Ala Leu Glu Pro Ser Gly Glu Ala Phe Asn Glu Leu Ser Leu Asp
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- Ile Asp Arg Val Leu Ala Glu Ser Glu Leu Ala Gly Ala Asp Arg Asn
- Ala His Asp Ala Val Leu Ser Tyr Thr Leu Gln Cys Ala Ile Lys Val 165 170 175
- Thr Thr Arg Asp Leu Ala Val Met Thr Ala Thr Leu Ala Ala Gly Gly
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- Leu Thr Leu Ser Val Met Ala Ser Ala Gly Met Tyr Asp Glu Ala Gly 210 215 220
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Trp 310	Trp				315				Gly	320					325	
ggt 112		cca	cag	ctg	cag	tac	atg	cca	gag	gaa	gaa	ggg	aca	gaa	aac	
Gly	Pro	Pro	Gln	. Leu 330		Tyr	Met	Pro	Glu 335		Glu	Gly	Thr	Glu 340	Asn	
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Ala	Leu	. Lev	ı Ala 20		a Thr	Let	ı Lev	Ala 25		. Cys	Thr	Pro	Thr 30		Val	

Glu Pro Ala Glu Thr Leu Thr Ala Leu Asp Pro Asp Ala Gly Pro Pro 40 Leu Pro Pro Asp Ser Ser Ile Glu Ala Pro Gly Glu Lys Glu Pro Ile 55 Val Glu Val Ile Glu Asn Trp Pro Gly Ser Leu Arg Pro Asp Asp Leu Thr Pro Glu Glu Arg Val Pro Gly Ile Val Asn Arg Gly Arg Ile Ile Val Gly Val Asp Gln Ser Gln Asn Leu Leu Ser Phe Arg Asp Pro Val Thr Gly Glu Leu Arg Gly Phe Glu Val Glu Leu Ala Arg Glu Ile Ser Arg Asp Ile Phe Gly Asp Pro Asn Lys Val Asp Phe Arg Phe Val Gly Ser Ser Asp Arg Leu Arg Ser Leu Asp Gln Gly Asp Val Asp Ile Val 150 Ile Arg Ser Val Thr Ile Thr Asp Glu Arg Ala Lys Leu Val Glu Phe 170 Ser Thr Pro Tyr Leu Arg Thr Gln Thr Arg Met Leu Thr Met Glu Ser Ser Gly Ile Thr Ser Ile Ala Asp Leu Pro Gly His Thr Ile Cys Val 200 Thr Asp Gly Ser Thr Ser Leu Gln Arg Ala Arg Thr Ile Ala Pro Glu 210 Ala Ser Ile Leu Lys Thr Arg Asn Trp Ser Asp Cys Leu Met Ala Leu 230 235 Gln Gln His Gln Ala Gln Val Ile Leu Gly Asp Asp Val Ile Leu Ser 250 Gly Ile Ala Ala Gln Asp Pro Tyr Thr Glu Ile Leu Asp Thr Ser Leu Asp Ser His Ser Tyr Gly Val Ala Ala Ala Ser Thr Thr Ala Glu Thr 280 Asp Ser Ser Gly Leu Ile Arg Gln Val Asn Tyr Thr Ile Glu Arg Ile Arg Thr Asp Arg Met Trp Trp Thr Met Phe Asp Asp Trp Phe Gly Pro Tyr Leu Trp Ser Tyr Gly Pro Pro Gln Leu Gln Tyr Met Pro Glu Glu 330 Glu Gly Thr Glu Asn Asp Glu Gly 340

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			atc gtg gca acc g Ile Val Ala Thr g 20										
			cag ggc aac cca g Gln Gly Asn Pro 2 35										
			gag att cag gcg a Glu Ile Gln Ala 1 50										
			acc gcc ggt gcc a Thr Ala Gly Ala a 65										
			gat ggt cag att a Asp Gly Gln Ile										
			ggg gtg atg ggc Gly Val Met Gly 1										
			atc ctt cca tcg of Ile Leu Pro Ser of 115										
	Leu Asp Ile G		ttc act gac aac Phe Thr Asp Asn 130										
gaa cgc cgc gag Glu Arg Arg Glu 135	aac ttt gat t Asn Phe Asp E 140	ttc atc gat ttc Phe Ile Asp Phe	ctc ttc gca ggt Leu Phe Ala Gly 145	gtg 547 Val									
			atc gat ccg gaa Ile Asp Pro Glu										
gcc tgt ggt ctc Ala Cys Gly Leu	acc gtt gct g Thr Val Ala V	gta cag cgc aca Val Gln Arg Thr 175	acc gtg gca gag Thr Val Ala Glu 180	acc 643 Thr									

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	_				_	•		gaa Glu 205		_	-			_		_	739
								gca Ala									787
								gaa Glu									835
	_		_	_				ggt Gly		-		_	_			_	883
								gcg Ala									931
								caa Gln 285									979
	gat 1032	• •	gcc	ctg	atc	aac	gaa	cag	сса	ctc	aac	taga	agcct	tc o	cagca	actaa	
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2/11115 1118

<212> PRT

<213> Corynebacterium glutamicum

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Gln Gly Asn Pro Asp Gly Trp Glu Gln Ile Val Pro Asp Pro Val Pro 35 40 45

Glu Ile Gln Ala Met Val Pro Glu Ala Leu Ala Gln Arg Gly Val Leu 50 55 60

Thr Ala Gly Ala Asn Pro Pro Phe Pro Pro Phe Glu Phe Lys Asp Ser 65 70 75 80

Asp Gly Gln Ile Ile Gly Val Glu Met Asp Leu Val Arg Ala Met Ala 85 90 95

Gly Val Met Gly Leu Glu Phe Ser Pro Gln Glu Gln Asp Phe Ser Leu 105 Ile Leu Pro Ser Val Gln Ala Gly Thr Leu Asp Ile Gly Ala Ser Gly 120 Phe Thr Asp Asn Glu Glu Arg Arg Glu Asn Phe Asp Phe Ile Asp Phe Leu Phe Ala Gly Val Gln Trp Ala Gln Ala Thr Asp Arg Glu Thr Pro 150 Ile Asp Pro Glu Asn Ala Cys Gly Leu Thr Val Ala Val Gln Arg Thr 170 165 Thr Val Ala Glu Thr Asp Asp Val Arg Pro Arg Ser Ala Gln Cys Glu 185 Ala Glu Gly Lys Glu Pro Ile Thr Ile Leu Ser Tyr Glu Thr Ala Asp 200 195 Thr Ala Ala Thr Ala Leu Ile Leu Gly Arg Ala Asp Ala Leu Ala Ala 215 Asp Ser Pro Val Ser Ala Trp Ala Ala Glu Arg Ser Glu Gly Arg Ile 235 230 Glu Val Val Gly Asp Met Tyr Leu Ala Ala Pro Phe Gly Phe Ala Phe 250 245 Pro Leu Glu Ser Asp Leu Thr Pro Ala Ala Ala Ala Phe Gln His 265 260 Leu Ile Asp Thr Gly Asp Tyr Gln Arg Ile Met Ala Gln Trp Gly Ile 280 Glu Glu Gly Leu Leu Asp Glu Ala Leu Ile Asn Glu Gln Pro Leu Asn 295 290

Pro	Asp	Asp	Ala	Gly 10	Thr	Trp	His	Asp	Ala 15	Asp	Ala	Ala	Phe	Gly 20	Phe	
	-			atc Ile		_									-	211
				gat Asp												259
				tac Tyr	-											307
				act Thr												355
				gag Glu 90												403
				gat Asp												451
				aag Lys												499
		120														
		tcc		gag Glu			acc					gcc				547
Val	Phe 135 cta	tcc Ser	Ser ctg		Lys	Lys 140 gat	acc Thr	Ile cgc	Leu	Glu	Met 145 gag	gcc Ala	Glu	Glu gtg	Met gac	547 595
Val aat Asn 150	Phe 135 cta Leu cag	tcc Ser gat Asp	Ser ctg Leu	Glu ggc	Lys ctt Leu 155 gag	Lys 140 gat Asp	acc Thr aag Lys	cgc Arg	Leu acc Thr	Glu att Ile 160 cac	Met 145 gag Glu gcg	gcc Ala cac His	Glu tac Tyr	Glu gtg Val	gac Asp 165	
Val  aat Asn 150 ttg Leu	Phe 135 cta Leu cag Gln	tcc Ser gat Asp tac Tyr	Ser ctg Leu gtg Val	Glu ggc Gly ccc Pro	Ctt Leu 155 gag Glu	Lys 140 gat Asp cca Pro	acc Thr aag Lys gat Asp	cgc Arg acc Thr	acc Thr ctt Leu 175	Glu att Ile 160 cac His	Met 145 gag Glu gcg Ala	gcc Ala cac His cag Gln	Glu tac Tyr att Ile	gtg Val tcc Ser 180	gac Asp 165 cgc Arg	595
Val  aat Asn 150 ttg Leu  ttg Leu  cag	Phe 135 cta Leu cag Glr gag Glu	tcc Ser gat Asp tac Tyr tca Ser	ctg Leu gtg Val ggc Gly 185	ggc Gly ccc Pro 170	Ctt Leu 155 gag Glu acc Thr	Lys 140 gat Asp cca Pro gca Ala	acc Thr aag Lys gat Asp aca Thr	cgc Arg acc Thr gtt Val 190	acc Thr ctt Leu 175 cgt Arg	Glu att Ile 160 cac His ccg Pro	Met 145 gag Glu gcg Ala ggc Gly	gcc Ala cac His cag Gln ggc Gly	tac Tyr att Ile aag Lys 195	gtg Val tcc Ser 180 ctg Leu	gac Asp 165 cgc Arg gaa Glu	595 643
val  aat Asn 150 ttg Leu  ttg Cag Gln ggt	Phe 135 cta Leu cag Gln gag Glu aag Lys	tcc Ser gat Asp tac Tyr tca Ser cgt Arg 200	ctg Leu gtg Val ggc Gly 185 tac Tyr	ggc Gly ccc Pro 170 tgc Cys	ctt Leu 155 gag Glu acc Thr	Lys 140 gat Asp cca Pro gca Ala	acc Thr aag Lys gat Asp aca Thr cag Gln 205	cgc Arg acc Thr gtt Val 190 ttc Phe cgc	acc Thr ctt Leu 175 cgt Arg cca Pro	Glu att Ile 160 cac His ccg Pro gta Val	Met 145 gag Glu gcg Ala ggc Gly cag Gln	gcc Ala cac His cag Gln ggc Gly aag Lys 210 gtg	tac Tyr att Ile aag Lys 195 gtc Val	gtg Val tcc Ser 180 ctg Leu gta Val	gac Asp 165 cgc Arg gaa Glu aag Lys	595 643 691
val  aat Asn 150 ttg Leu  ttg Cag Gln  ggt Gly agc	Phe 135 cta Leu cag Glr gag Glu aag Lys 215 gtc	tcc Ser gat Asp tac Tyr tca Ser cgt Arg 200 gag Glu	ctg Leu gtg Val ggc Gly 185 tac Tyr cag Gln	ggc Gly ccc Pro 170 tgc Cys ttc Phe	Ctt Leu 155 gag Glu acc Thr aag Lys	Lys 140 gat Asp cca Pro gca Ala cct Pro ttc Phe 220	acc Thr aag Lys gat Asp aca Thr cag Gln 205 gat Asp	cgc Arg acc Thr gtt Val 190 ttc Phe cgc Arg gac	acc Thr ctt Leu 175 cgt Arg cca Pro att Ile	Glu att Ile 160 cac His ccg Pro gta Val gcc Ala acc	Met 145 gag Glu gcg Ala ggc Gly cag Gln 225 gta	gcc Ala cac His cag Gln ggc Gly aag Lys 210 gtg Val	tac Tyr att Ile aag Lys 195 gtc Val ttg Leu	gtg Val tcc Ser 180 ctg Leu gta Val	gac Asp 165 cgc Arg gaa Glu aag Lys gat Asp	<ul><li>595</li><li>643</li><li>691</li><li>739</li></ul>

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tcg gag gtc gat Ser Glu Val Asp 280	gtg gct gcg Val Ala Ala	gag tcc Glu Ser 285	gcc gct gcg Ala Ala Ala	att ggc Ile Gly 290	gct gag 979 Ala Glu
cac atc gtg aag	att gtc tcg	cct gag	gaa tac gcc	aac gcg	att cct
His Ile Val Lys 295	: Ile Val Ser 300		Glu Tyr Ala 305	Asn Ala	Ile Pro
aag atc atg tgg	g tac ttg gat	gat cct	gta gct gac	cca tca	ttg gtc
Lys Ile Met Trp 310	Tyr Leu Asp 315	Asp Pro	Val Ala Asp 320	Pro Ser	Leu Val 325
ccg ctg tac ttc	gtg gca gcg	gaa gca	cgt aag cac	gtc aag	gtt g <b>tg</b>
Pro Leu Tyr Phe	e Val Ala Ala 330	Glu Ala	Arg Lys His	Val Lys	Val Val 340
ctg tot ggc gag	g ggc gca gat	gag ctg	ttc ggt gga	tac acc	att tac
Leu Ser Gly Glv		Glu Leu 350		Tyr Thr 355	Ile Tyr
aag gag ccg cta 1219	a teg ett get	cca ttt	gag aag ato	cct tcc	cca cta
Lys Glu Pro Let 360	u Ser Leu Ala	a Pro Phe 365	Glu Lys Ile	Pro Ser 370	Pro Leu
cgt aaa ggc ct	g gga aag cto	agc aag	gtt ctg cca	gac ggc	atg aag
Arg Lys Gly Le 375	u Gly Lys Lev 380		Val Leu Pro 385	Asp Gly	Met Lys
ggc aag tcc ct 1315	t ctt gag cg	ggc tcc	atg acc atg	g gaa gag	cgc tac
Gly Lys Ser Le 390	u Leu Glu Arg 395	g Gly Ser	Met Thr Met 400	: Glu Glu	Arg Tyr 405
tac ggc aac gc 1363	t cgc tcc tte	c aat tto	gag cag ato	g caa cgc	gtt att
Tyr Gly Asn Al	a Arg Ser Pho 410	e Asn Phe	e Glu Gln Met 415	Gln Arg	Val Ile 420
cca tgg gca aa	g cgc gaa tg	g gac cac	cgc gaa gto	c act gcg	ccg atc
1411 Pro Trp Ala Ly 42		p Asp His 430		1 Thr Ala 435	Pro Ile
tac gca cag to 1459	c cgc aac tt	t gat cca	a gta gcc cg	c atg caa	cac ctg
Tyr Ala Gln Se 440	r Arg Asn Ph	e Asp Pro 445	o Val Ala Ar	g Met Gln 450	His Leu

gat ctg ttc acc tgg atg cgc ggc gac atc ctg gtc aag gct gac aag 1507 Asp Leu Phe Thr Trp Met Arg Gly Asp Ile Leu Val Lys Ala Asp Lys 460 atc aac atg gcg aac tcc ctt gag ctg cga gtt cca ttc ttg gat aag 1555 Ile Asn Met Ala Asn Ser Leu Glu Leu Arg Val Pro Phe Leu Asp Lys 470 475 480 gaa gtt ttc aag gtt gca gag acc att cct tac gac ctg aag att gcc 1603 Glu Val Phe Lys Val Ala Glu Thr Ile Pro Tyr Asp Leu Lys Ile Ala 495 490 aac ggt acc acc aag tac gcg ctg cgc agg gca ctc gag cag att gtt 1651 Asn Gly Thr Thr Lys Tyr Ala Leu Arg Arg Ala Leu Glu Gln Ile Val 505 510 ccg cct cac gtt ttg cac cgc aag aag ctg ggc ttc cct gtt ccc atg 1699 Pro Pro His Val Leu His Arg Lys Lys Leu Gly Phe Pro Val Pro Met 525 cgc cac tgg ctt gcc ggc gat gag ctg ttc ggt tgg gcg cag gac acc 1747 Arg His Trp Leu Ala Gly Asp Glu Leu Phe Gly Trp Ala Gln Asp Thr 540 545 atc aag gaa too ggt act gaa gat atc tto aac aag cag got gtg otg 1795 Ile Lys Glu Ser Gly Thr Glu Asp Ile Phe Asn Lys Gln Ala Val Leu 555 560 gat atg etg aac gag cac ege gat gge gtg tea gat cat tee egt ega 1843 Asp Met Leu Asn Glu His Arg Asp Gly Val Ser Asp His Ser Arg Arg 570 575 580 ctg tgg act gtt ctg tca ttt atg gtg tgg cac ggc att ttt gtg gaa 1891 Trp His Gly lie Phe Val Glu vai 585 590 595 aac cgc att gat cca cag att gag gac cgc tcc tac cca gtc gag ctt 1939 Asn Arg Ile Asp Pro Gln Ile Glu Asp Arg Ser Tyr Pro Val Glu Leu 600 taagtettaa ageetaaace eec 1962

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<211> 613

<212> PRT

<213> Corynebacterium glutamicum

<400> 110

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His	Gln	Pro 35	Leu	Arg	Trp	Gly	Pro 40	Ala	Asp	Glu	Pro	Asp 45	Arg	Tyr	Ala
Met	Thr 50	Phe	Asn	Gly	Glu	Ile 55	Туг	Asn	Tyr	Val	Glu 60	Leu	Arg	Lys	Glu
Leu 65	Ser	Asp	Leu	Gly	Tyr 70	Ala	Phe	Asn	Thr	Ser 75	Gly	Asp	Gly	Glu	Pro 80
Ile	Val	Val	Gly	Phe 85	His	His	Trp	Gly	Glu 90	Ser	Val	Val	Glu	His 95	Leu
Arg	Gly	Met	Phe 100	Gly	Ile	Ala	Ile	Trp 105	Asp	Thr	Lys	Glu	Lys 110	Ser	Leu
Phe	Leu	Ala 115	Arg	Asp	Gln	Phe	Gly 120	Ile	Lys	Pro	Leu	Phe 125	Tyr	Ala	Thr
Thr	Glu 130	His	Gly	Thr	Val	Phe 135	Ser	Ser	Glu	Lys	Lys 140	Thr	Ile	Leu	Glu
Met 145	Ala	Glu	Glu	Met	Asn 150	Leu	Asp	Leu	Gly	Leu 155	Asp	Lys	Arg	Thr	Ile 160
Glu	His	Tyr	Val	Asp 165	Leu	Gln	Tyr	Val	Pro 170	Glu	Pro	Asp	Thr	Leu 175	His
Ala	Gln	Ile	Ser 180	Arg	Leu	Glu	Ser	Gly 185	Cys	Thr	Ala	Thr	Val 190	Arg	Pro
Gly	Gly	Lys 195		Glu	Gln	Lys	Arg 200	Tyr	Phe	Lys	Pro	Gln 205	Phe	Pro	Val
Gln	Lys 210		Val	Lys	Gly	Lys 215		Gln	Asp	Leu	Phe 220	Asp	Arg	Ile	Ala
Gln 225		Leu	Glu	Asp	Ser 230	Val	Glu	Lys	His	Met 235	Arg	Ala	Asp	Val	Thr 240
Val	Gly	Ser	Phe	Leu 245		Gly	Gly	Ile	Asp 250	Ser	Thr	Ala	Ile	Ala 255	Ala
Leu	Ala	Lys	260		Asn	Pro	Asp	Leu 265	Leu	Thr	Phe	Thr	Thr 270	Gly	Phe
Glu	a Arg	g Glu 275		Tyr	Ser	Glu	Val 280		Val	Ala	Ala	Glu 285	Ser	Ala	Ala
Ala	11e 290		/ Ala	Glu	His	Ile 295		Lys	Ile	Val	Ser 300		Glu	Glu	Tyr
Ala 305		n Alá	a Ile	e Pro	) Lys 310		Met	Trp	Tyr	Leu 315	Asp	Asp	Pro	Val	Ala 320
Asp	Pro	Sei	. Leu	val 325		Leu	. Tyr	Phe	val 330		Ala	Glu	Ala	Arg 335	Lys

His Val Lys Val Val Leu Ser Gly Glu Gly Ala Asp Glu Leu Phe Gly 340 345 Gly Tyr Thr Ile Tyr Lys Glu Pro Leu Ser Leu Ala Pro Phe Glu Lys 360 Ile Pro Ser Pro Leu Arg Lys Gly Leu Gly Lys Leu Ser Lys Val Leu 380 370 375 Pro Asp Gly Met Lys Gly Lys Ser Leu Leu Glu Arg Gly Ser Met Thr 390 395 Met Glu Glu Arg Tyr Tyr Gly Asn Ala Arg Ser Phe Asn Phe Glu Gln 405 410 Met Gln Arg Val Ile Pro Trp Ala Lys Arg Glu Trp Asp His Arg Glu Val Thr Ala Pro Ile Tyr Ala Gln Ser Arg Asn Phe Asp Pro Val Ala 440 445 Arg Met Gln His Leu Asp Leu Phe Thr Trp Met Arg Gly Asp Ile Leu 455 Val Lys Ala Asp Lys Ile Asn Met Ala Asn Ser Leu Glu Leu Arg Val 470 475 Pro Phe Leu Asp Lys Glu Val Phe Lys Val Ala Glu Thr Ile Pro Tyr 490 Asp Leu Lys Ile Ala Asn Gly Thr Thr Lys Tyr Ala Leu Arg Arg Ala 500 505 Leu Glu Gln Ile Val Pro Pro His Val Leu His Arg Lys Lys Leu Gly 520 Phe Pro Val Pro Met Arg His Trp Leu Ala Gly Asp Glu Leu Phe Gly 530 535 540 Trp Ala Gln Asp Thr Ile Lys Glu Ser Gly Thr Glu Asp Ile Phe Asn 545 550 555 Lys Gln Ala Val Leu Asp Met Leu Asn Glu His Arg Asp Gly Val Ser 565 570 Asp His Ser Arg Arg Leu Trp Thr Val Leu Ser Phe Met Val Trp His 580 585 Gly The Phe Val Glu Asn Arg Ile Asp Pro Gln Ile Glu Asp Arg Ser 600 Tyr Pro Val Glu Leu 610 <210> 111 <211> 1284 <212> DNA <213> Corynebacterium glutamicum

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        200
                                                                   787
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Lys His Val Ser Val Ala Lys Leu Pro Gly Met Trp Asp Arg Thr Val
                        220
                                            225
acg gtg tcg tcg gcg gcg aaa acg ttc aat gtg act ggt tgg aag acg
                                                                   835
Thr Val Ser Ser Ala Ala Lys Thr Phe Asn Val Thr Gly Trp Lys Thr
                                        240
                    235
                                                                   883
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Gly Trp Ala Leu Ala Pro Glu Pro Leu Leu Glu Ala Val Leu Lys Ala
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                                    255
aug cag tit atg tot tat gtg ggg got aca cot tit cag cog got gtg
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Lys Gln Phe Met Ser Tyr Val Gly Ala Thr Pro Phe Gln Pro Ala Val
            265
                                270
                                                                   979
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Ala His Ala Ile Glu His Glu Gln Lys Trp Val Ser Lys Met Ser Lys
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        280
ggg ctt gag ctc aag cgg gat att ttg cgt act gcg tta gat aag gcg
1027
Gly Leu Glu Leu Lys Arg Asp Ile Leu Arg Thr Ala Leu Asp Lys Ala
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310
att ggg gat cgt gat ggt gcg gag ttc tgt ttt gag ttg att gag aag
1123
Ile Gly Asp Arg Asp Gly Ala Glu Phe Cys Phe Glu Leu Ile Glu Lys
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gtt ggg gtg gcg att ccg gtg cag gcg ttt gtg gat cat ccg aag
1171
Val Gly Val Ala Ala Ile Pro Val Gln Ala Phe Val Asp His Pro Lys
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tagtttgaac aggttgttgg ggg 1284

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<211> 387

<212> PRT

<213> Corynebacterium glutamicum

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Lys Thr Thr Val Gln Lys Lys Phe Arg Ile Glu Ser Asp Leu Leu Gly 50

Glu Leu Gln Ile Pro Ser His Ala Tyr Tyr Gly Val His Thr Leu Arg 70

Ala Val Asp Asn Phe Gln Ile Ser Arg Thr Thr Ile Asn His Val Pro 85 90 95

Asp Phe Ile Arg Gly Met Val Gln Val Lys Lys Ala Ala Ala Leu Ala 100 105 110

Asn Arg Arg Leu His Thr Leu Pro Ala Gln Lys Ala Glu Ala Ile Val 120 Trp Ala Cys Asp Gln Ile Leu Ile Glu Glu Arg Cys Met Asp Gln Phe 135 Pro Ile Asp Val Phe Gln Gly Gly Ala Gly Thr Ser Leu Asn Met Asn Thr Asn Glu Val Val Ala Asn Leu Ala Leu Glu Phe Leu Gly His Glu 170 Lys Gly Glu Tyr His Ile Leu His Pro Met Asp Asp Val Asn Met Ser 185 180 Gln. Ser Thr Asn Asp Ser Tyr Pro Thr Gly Phe Arg Leu Gly Ile Tyr Ala Gly Leu Gln Thr Leu Ile Ala Glu Ile Asp Glu Leu Gln Val Ala 215 Phe Arg His Lys Gly Asn Glu Phe Val Asp Ile Ile Lys Met Gly Arg 235 230 Thr Gln Leu Gln Asp Ala Val Pro Met Ser Leu Gly Glu Glu Phe Arg 245 Ala Phe Ala His Asn Leu Ala Glu Glu Gln Thr Val Leu Arg Glu Ala 265 Ala Asn Arg Leu Leu Glu Val Asn Leu Gly Ala Thr Ala Ile Gly Thr 280 Gly Val Asn Thr Pro Ala Gly Tyr Arg His Gln Val Val Ala Ala Leu Ser Glu Val Thr Gly Leu Glu Leu Lys Ser Ala Arg Asp Leu Ile Glu 310 305 Ala Thr Ser Asp Thr Gly Ala Tyr Val His Ala His Ser Ala Ile Lys 330 Arg Ala Ala Met Lys Leu Ser Lys Ile Cys Asn Asp Leu Arg Leu Leu Ser Ser Gly Pro Arg Ala Gly Leu Asn Glu Ile Asn Leu Pro Pro Arg 360 Gln Ala Gly Ser Ser Ile Met Pro Ala Lys Val Asn Pro Val Ile Pro 375 370 Glu Val Val Asn Gln Val Cys Phe Lys Val Phe Gly Asn Asp Leu Thr 395 390 Val Thr Met Ala Ala Glu Ala Gly Gln Leu Gln Leu Asn Val Met Glu 405 Pro Val Ile Gly Glu Ser Leu Phe Gln Ser Leu Arg Ile Leu Gly Asn 425 Ala Ala Lys Thr Leu Arg Glu Lys Cys Val Val Gly Ile Thr Ala Asn

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tgg cac acc Trp His Thr			Ala P							691
cca gag cgc Pro Glu Arg 200	Pro Asp									739
gtc gaa atc Val Glu Ile 215										787
gct gcc atc Ala Ala Ile 230										835
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Cys Thr Ser Asp Ala Asn Gly His Leu Leu Pro Thr Val Ser Gly Ala 35 40 45

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Phe Glu Ile His Glu Ile Asn Arg Leu Asp Ser Ser Ser Met Thr Phe 65 70 75 80

Glu Asp Leu Asp Ser Ile Ile Ala Thr Val His Lys Val Leu Glu Asp 85 90 95

Pro Asp Val Val Gly Val Val Val Thr His Gly Thr Asp Ser Met Glu 100 105 110

Glu Ser Ala Ile Ala Val Asp Thr Phe Leu Asp Asp Pro Arg Pro Val

Ile Phe Thr Gly Ala Gln Lys Pro Phe Asp His Pro Glu Ala Asp Gly 130 135 140

Pro Asn Asn Leu Phe Glu Ala Cys Leu Ile Ala Ser Asp Pro Ser Ala 145 150 155 160

Arg Gly Ile Gly Ala Leu Ile Val Phe Gly His Ala Val Ile Pro Ala 165 170 175

Arg Gly Cys Val Lys Trp His Thr Ser Asp Glu Leu Ala Phe Ala Thr 180 185 190

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Val Glu Ala Met Gly Ser Gly Asn Val Gly Ser Arg Met Gly Asp Ala 245 250 255

Leu Gly Lys Ala Leu Asp Ala Gly Ile Pro Val Val Met Ser Thr Arg
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Val Pro Arg Gly Glu Val Ser Gly Val Tyr Gly Gly Ala Gly Gly Gly 275 280 285

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Ile Ala Ala His Gly Ala Asp Ala Phe Gly Val Ala Thr Leu Ala Glu  $50 \hspace{1cm} 55 \hspace{1cm} 60$ 

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Asp Ala Glu His Ile Arg Val Ser Ile Lys Ile Asp Ser Gly Leu His
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Arg Ser Gly Val Asp Glu Gln Glu Trp Glu Gly Val Phe Ser Ala Leu 130 140

Ala Ala Ala Pro His Ile Glu Val Thr Gly Met Phe Thr His Leu Ala 145 150 155 160

Cys Ala Asp Glu Pro Glu Asn Pro Glu Thr Asp Arg Gln Ile Ile Ala 165 170 175

Phe Arg Arg Ala Leu Ala Leu Ala Arg Lys His Gly Leu Glu Cys Pro 180 185 190

Val Asn His Val Cys Asn Ser Pro Ala Phe Leu Thr Arg Ser Asp Leu 195 200 205

His Met Glu Met Val Arg Pro Gly Leu Ala Phe Tyr Gly Leu Glu Pro 210 215 220

Val Ala Gly Leu Glu His Gly Leu Lys Pro Ala Met Thr Trp Glu Ala 225 230 235 240

Lys Val Ser Val Val Lys Gln Ile Glu Ala Gly Gln Gly Thr Ser Tyr 245 250 255

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tcat tcga	ggga	att o	ctc	cgaa	ac at	cgc	gaata gag	tto cgc	cctac	act	atg Met 1 gcc	atg Met aac	att Ile att	gat Asp	aca Thr 5 agg	
tcat tcga	ggga atoto got	ggc ogtt Val	ctc Leu	att Ile 10	gac Asp	cgc Arg	gag Glu cat	cgc Arg	tta Leu 15	act Thr	atg Met 1 gcc Ala	atg Met aac Asn	att Ile att Ile	gat Asp tcc Ser 20	aca Thr 5 agg Arg	115
tcat tcga cct Pro atg Met	ggga atcto gct Ala	ggc ogtt Val gct Ala	ctc Leu Cac His 25	att Ile 10 gcc Ala	gac Asp ggt Gly	cgc Arg gcc Ala	gag Glu cat His	cgc Arg gag Glu 30	tta Leu 15 att Ile	act Thr gcc Ala	atg Met 1 gcc Ala ctg Leu	atg Met aac Asn cgt Arg	att Ile att Ile ccg Pro 35	gat Asp tcc Ser 20 cat His	aca Thr 5 agg Arg Val	115
tcat tcga cct Pro atg Met aaa Lys	gct Ala gca Ala acg	gtt Val  gct Ala  cac His 40	ctc Leu Cac His 25 aaa Lys	att Ile 10 gcc Ala atc Ile	gac Asp ggt Gly att Ile	cgc Arg gcc Ala gaa Glu gca	gag Glu cat His att Ile 45	cgc Arg gag Glu 30 gcg Ala	tta Leu 15 att Ile cag Gln	act Thr gcc Ala atg Met	atg Met 1 gcc Ala ctg Leu cag Gln	atg Met aac Asn cgt Arg gtc Val 50 gaa	att Ile att Ile ccg Pro 35 gac Asp	gat Asp tcc Ser 20 cat His	aca Thr 5 agg Arg Gtg Val ggt Gly	115 163 211
tcat tcga cct Pro atg Met aaa Lys gcc Ala	gct Ala gca Ala acg Thr	gtt Val  gct Ala  cac His 40  ggg	ctc Leu cac His 25 aaa Lys atc Ile	att Ile 10 gcc Ala atc Ile acc Thr	gac Asp ggt Gly att Ile tgc Cys	cgc Arg gcc Ala gaa Glu gca Ala 60	gag Glu cat His att Ile 45 acc Thr	cgc Arg gag Glu 30 gcg Ala att	tta Leu 15 att Ile cag Gln ggc Gly	act Thr gcc Ala atg Met gag Glu	atg Met 1 gcc Ala ctg Leu cag Gln gcg Ala 65	atg Met aac Asn cgt Arg gtc Val 50 gaa Glu	att Ile att Ile ccg Pro 35 gac Asp att Ile tat	gat Asp tcc Ser 20 cat His gcc Ala	aca Thr 5 agg Arg  gtg Val  ggt Gly  gcc Ala acc	115 163 211 259

ggc (	gtg Val	gat Asp	tcg Ser 105	gta Val	gag Glu	atg Met	gca Ala	cag Gln 110	gcg Ala	acg Thr	gcg Ala	ggt Gly	ttg Leu 115	cgg Arg	gaa Glu	451
gat Asp	atc Ile	aag Lys 120	gct Ala	ctg Leu	att Ile	gaa Glu	gtg Val 125	gat Asp	tcg Ser	gga Gly	cat His	cgt Arg 130	aga Arg	agt Ser	gga Gly	499
gtc Val	acg Thr 135	gcg Ala	act Thr	gct Ala	tca Ser	gaa Glu 140	ttg Leu	agt Ser	cag Gln	atc Ile	cgc Arg 145	gag Glu	gcg Ala	ctg Leu	ggc Gly	547
agc Ser 150	agg Arg	tat Tyr	gca Ala	gga Gly	gtg Val 155	ttt Phe	act Thr	ttt Phe	cct Pro	ggg Gly 160	cat His	tct Ser	tat Tyr	ggc Gly	ccg Pro 165	595
gga Gly	aat Asn	ggt Gly	gag Glu	cag Gln 170	gca Ala	gca Ala	gct Ala	gat Asp	gag Glu 175	ctt Leu	cag Gln	gct Ala	cta Leu	aac Asn 180	aac Asn	643
agc Ser	gtc Val	cag Gln	cga Arg 185	ctt Leu	gct Ala	ggc Gly	ggc Gly	ctg Leu 190	act Thr	tct Ser	ggc Gly	ggt Gly	tcc Ser 195	tcg Ser	ccg Pro	691
tct Ser	gcg Ala	cag Gln 200	ttt Phe	aca Thr	gac Asp	gca Ala	atc Ile 205	gat Asp	gag Glu	atg Met	cga Arg	cca Pro 210	ggc Gly	gtg Val	tat Tyr	739
gtg Val	ttt Phe 215	Asn	gat Asp	tcc Ser	cag Gln	cag Gln 220	atc Ile	acc Thr	tcg Ser	gga Gly	gca Ala 225	tgc Cys	act Thr	gag Glu	aag Lys	787
cag Gln 230	gtg Val	gca Ala	atg Met	acg Thr	gtg Val 235	Leu	tct Ser	act Thr	gtg Val	gtc Val 240	agc Ser	cga Arg	aat Asn	gtg Val	tca Ser 245	835
gat Asp	cgt Arg	cgg Arg	atc Ile	att Ile 250	Leu	gat Asp	gcg Ala	gga Gly	tcc Ser 255	Lys	atc Ile	ctc Leu	agc Ser	act Thr 260	gat Asp	883
Lys	Pro	Ala	265	Ile	e Asp	Gly	Asn	Gly 270	Phe	. Val	. Leu	Gly	275		GIU	931
gcc Ala	cga Arg	atc ; Ile 280	Ser	gct Ala	ttg Lei	tcg Ser	gag Glu 285	His	cac His	gca Ala	acc Thr	att Ile 290	Phe	tgg Trp	cca Pro	979
gat 102		a gto	g cta	ctt	cca	a gta	ato	ggg	gag	g cag	gcto	aac	ato	gtg	CCC	
Asp	295	5				300	)				305	)			Pro	
107	5														cgg	
Asn 310	n Hi:				319	5				320	0				325	
gaa 112		c gat	t ggd	c act	t tto	c cg1	t acc	c tgg	g aaq	g gta	a gtt	ged	c cgo	ggc	aga	

Glu Ala Asp Gly Thr Phe Arg Thr Trp Lys Val Val Ala Arg Gly Arg 330 335 340

aac aat tagggaaacc tettgacett cac 1152 Asn Asn

<210> 132

<211> 343

<212> PRT

<213> Corynebacterium glutamicum

<400> 132

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Ala Asn Ile Ser Arg Met Ala Ala His Ala Gly Ala His Glu Ile Ala 20 25 30

Leu Arg Pro His Val Lys Thr His Lys Ile Ile Glu Ile Ala Gln Met 35 40 45

Gln Val Asp Ala Gly Ala Arg Gly Ile Thr Cys Ala Thr Ile Gly Glu 50 60

Ala Glu Ile Phe Ala Gly Ala Gly Phe Thr Asp Ile Phe Ile Ala Tyr 65 70 75 80

Pro Leu Tyr Leu Thr Asp His Ala Val Gln Arg Leu Asn Ala Ile Pro 85 90 95

Gly Glu Ile Ser Ile Gly Val Asp Ser Val Glu Met Ala Gln Ala Thr 100 105 110

Ala Gly Leu Arg Glu Asp Ile Lys Ala Leu Ile Glu Val Asp Ser Gly
115 120 125

His Arg Arg Ser Gly Val Thr Ala Thr Ala Ser Glu Leu Ser Gln Ile 130 135 140

His Ser Tyr Gly Pro Gly Asn Gly Glu Gln Ala Ala Asp Glu Leu 165 170 175

Gln Ala Leu Asn Asn Ser Val Gln Arg Leu Ala Gly Gly Leu Thr Ser

Gly Gly Ser Ser Pro Ser Ala Gln Phe Thr Asp Ala Ile Asp Glu Met
195 200 205

Arg Pro Gly Val Tyr Val Phe Asn Asp Ser Gln Gln Ile Thr Ser Gly 210 215 220

Ala Cys Thr Glu Lys Gln Val Ala Met Thr Val Leu Ser Thr Val Val 225 230 235 240

Ser Arg Asn Val Ser Asp Arg Ile Ile Leu Asp Ala Gly Ser Lys

				245					250					255		
Ile	Leu	Ser	Thr 260	Asp	Lys	Pro	Ala	Trp 265	Ile	Asp	Gly	Asn	Gly 270	Phe	Val	
Leu	Gly	Asn 275	Pro	Glu	Ala	Arg	Ile 280	Ser	Ala	Leu	Ser	Glu 285	His	His	Ala	
Thr	Ile 290	Phe	Trp	Pro	Asp	Lys 295	Val	Leu	Leu	Pro	Val 300	Ile	Gly	Glu	Gln	
Leu 305	Asn	Ile	Val	Pro	Asn 310	His	Ala	Cys	Asn	Val 315	Ile	Asn	Leu	Val	Asp 320	
Glu	Val	Tyr	Val	Arg 325	Glu	Ala	Asp	Gly	Thr 330	Phe	Arg	Thr	Trp	Lys 335	Val	
Val	Ala	Arg	Gly 340	Arg	Asn	Asn										
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	3> R	XA02														
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<22 <40 aag	0> 1 aagt	33 gat	536 cacg	cgaa							atg	gac Asp	aac	ttc	gatta gcc Ala 5	it 60 115
<222 <40 aag	0> 1 aagt tatt	33 gat gcc	536 cacg ttgc	cgaa ttca gct	ga t gct Ala	ctcg gaa	acga	a tt	ccga gcg	taag gaa Glu	atg Met 1	gac Asp	aac Asn gct	ttc Phe	gcc Ala 5 gtg Val	
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<222 <40 aag gcg ctg Let ttg Let Asg	0> 1 aagt tatt g ctg i Leu g gtg i Val	33 gat gcc cgt Arg ttt Phe cag Glr 40 a gcc	cacg ttgc gat Asp ccg Pro 25 gcg Ala	cgaa ttca gct Ala 10 gag Glu	ga t gct Ala ggcg Ala ggag Glu	gaa Glu act Thr cto Leu	acga aaaa Lys teg Ser Ser Asp 45	a tt gct Ala gcaa Gln 30 Gly	gcga Ala 15 ago Ser Glu	gaa Glu ttt Phe	atg Met 1 cag Glr ggt Gly	gac Asp ggg Gly acg Thr 50	aacc Asn gct Alaa gct Alaa gct Gly 35 Gly 35 Gly 35 ata	tto Phe cgg Arg 20 a agg V Arg	gcc Ala 5 gtg Val	115 163 211
<222 <400 aag gcg ctc Let ttc Let ttc Let Asg	0> 1 aagt tatt g ctg g tg g Val t actt t tes t gcf o Ala	33 gat gcc cgt Arg ttt Phe cag Glr 40 a gcc	cacg ttgc gat Asp ccg Pro 25 g gcg n Ala	cgaa ttca gct Ala 10 gag Glu gag Glu	ga t gct Ala g gcc n Ala g gaç n Ctç n Len	gaaa. Glu acti Thr	acga aaa Lys tegat Ser 45 Quit Vai	a tt gct Ala Gcaa Glr 30 Gly Val	gcga 15 ago Gluca a aaa	taagg	atg Met 1 cag Gln t ggt t Gly c tcc Ser t gcc 1 Ala 65	gac Asp ggg Gly acg Thr 50	aacc Asr gct gct as gct	cgg Arg Arg Arg Arg Arg Phe	gcc Ala 5 gtg Val ctt Leu Acga Arg	<ul><li>115</li><li>163</li><li>211</li><li>259</li></ul>

			gac Asp						-					451
			ctg Leu											499
			tac Tyr											547
_	_		gca Ala	_				-						595
			tta Leu 170											643
_	_		tgc Cys	-		_		_				_		691
_	_		gat Asp	_	_				-					739
			cca Pro											787
			atc Ile											835
			cct Pro 250			taad	ccac	igt (	ctaaq	ggaal	cc a	ct		879

<210> 134

<211> 252

<212> PRT

<213> Corynebacterium glutamicum

<400> 134

Met Asp Asn Phe Ala Leu Leu Arg Asp Ala Ala Glu Lys Ala Ala Glu  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Gin Gly Ala Arg Val Leu Val Phe Pro Glu Ala Thr Ser Gln Ser Phe 20 25 30

Gly Thr Gly Arg Leu Asp Thr Gln Ala Glu Glu Leu Asp Gly Glu Phe

Ser Thr Ala Val Arg Lys Leu Ala Asp Glu Leu Asp Val Val Ile Val 50 55 60

Ala Gly Met Phe Thr Pro Ala Asp Thr Val Gln Arg Gly Glu Lys Thr

65					70					75					80	
Ile	Ser	Arg	Val	Asn 85	Asn	Thr	Val	Leu	Ile 90	Ser	Gly	Ala	Gly	Leu 95	His	
Gln	Gly	Tyr	Asn 100	Lys	Ile	His	Thr	Tyr 105	Asp	Ala	Phe	Gly	Tyr 110	Arg	Glu	
Ser	Asp	Thr 115	Val	Lys	Pro	Gly	Asp 120	Glu	Leu	Val	Val	Phe 125	Glu	Val	Asp	
Asp	Ile 130	Lys	Phe	Gly	Val	Ala 135	Thr	Cys	Tyr	Asp	Ile 140	Arg	Phe	Pro	Glu	
Gln 145	Phe	Lys	Asp	Leu	Ala 150	Arg	Asn	Gly	Ala	Gln 155	Ile	Ile	Val	Val	Pro 160	
Thr	Ser	Trp	Gln	Asp 165	Gly	Pro	Gly	Lys	Leu 170	Glu	Gln	Trp	Glu	Val 175	Leu	
Pro	Arg	Ala	Arg 180	Ala	Leu	Asp	Ser	Thr 185	Cys	Trp	Ile	Val	Ala 190	Cys	Gly	
Gln	Ala	Arg 195	Leu	Pro	Glu	Glu	Leu 200	Arg	Asp	Glu	Arg	Lys 205	Gly	Pro	Thr	
Gly	11e 210	Gly	His	Ser	Met	Val 215	Thr	Asn	Pro	His	Gly 220	Glu	Val	Ile	Ala	
Ser 225	Ala	Gly	Tyr	Glu	Pro 230	Glu	Met	Leu	Ile	Ala 235	Asp	Ile	Asp	Val	Ser 240	
Gly	Leu	Ala	Lys	Ile 245	Arg	Glu	Ala	Leu	Pro 250	Val	Leu					
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caa	caatt	ca (	atta	gcaga	ag ca	attta	aagg	a ati	ttac	acac	_		gaa Glu		_	115
	atc Ile	_				-										163
	acc Thr															211

_		_	-	_				atc Ile					-		-	259
	_	_	-	_				aac Asn				_	_	_		307
_						_		ctg Leu	_		-	_	_			355
_						_		cac His	-	-	-	_		-	-	403
_								gaa Glu 110			_		-			451
			_					ttc Phe				_				499
	_				_			cca Pro	_		_	_				547
_	_					-	_	gtg Val	_	_						595
		_	-			_	_	att Ile	_	_				_	-	643
								cag Gln 190			_	-	_			691
								cag Gln								739
								tct Ser								787
								gag Glu								835
-								aag Lys			_	-		-		883
								gat Asp 270								931
ggc	gct	gcc	ggt	gag	cgt	tgc	atg	gct	gtt	tct	gtg	gtc	ttg	gct	att	979

Gly Ala Ala Gly Glu Arg Cys Met Ala Val Ser Val Val Leu Ala Ile 280 285 gaa tot gtt god gad gag oto att gag aag atd aag gag ogd atd gad 1027 Glu Ser Val Ala Asp Glu Leu Ile Glu Lys Ile Lys Glu Arg Ile Asp 300 acc ctg cgc atc ggc aac ggt gcc ggc gac gag cag ggc gag ccg cac 1075 Thr Leu Arg Ile Gly Asn Gly Ala Gly Asp Glu Gln Gly Glu Pro His 310 315 320 ctg ggc cca cta atc acc gac gtc cac cgc gac aag gtc gct tct tat 1123 Leu Gly Pro Leu Ile Thr Asp Val His Arg Asp Lys Val Ala Ser Tyr 335 gto gac atc gct gag gcc gac ggc gcc aag atc atc gtg gac ggg cgt 1171 Val Asp Ile Ala Glu Ala Asp Gly Ala Lys Ile Ile Val Asp Gly Arg 350 aac tgc gcc gta gac ggg cac gag gag ggc ttc ttc ttc ggc cct acg Asn Cys Ala Val Asp Gly His Glu Glu Gly Phe Phe Gly Pro Thr ctt atc gac gac atc cca ctc acg ttc cgc gcc tac acc gaa gaa atc Leu Ile Asp Asp Ile Pro Leu Thr Phe Arg Ala Tyr Thr Glu Glu Ile 380 385 ttc ggc ccg gtc ctc tct gtc gtt cgt gtc gca tcc ttc gac gag gca 1315 Phe Gly Pro Val Leu Ser Val Val Arg Val Ala Ser Phe Asp Glu Ala 395 att gag ctg atc aac tcc ggt gaa ttc ggc aac gga acc gca atc ttc 1363 Ile Glu Leu Ile Asn Ser Gly Glu Phe Gly Asn Gly Thr Ala Ile Phe 410 415 ace aac gat ggt gga geg gea ege ege tte eag eat gag ate gaa gtg 1411 Thr Asn Asp Gly Gly Ala Ala Arg Arg Phe Gln His Glu Ile Glu Val 430 ggc atg atc ggc atc aac gta cca atc cca gtg cct gtt gcg tac cac Gly Met Ile Gly Ile Asn Val Pro Ile Pro Val Pro Val Ala Tyr His 440 445 450 tcc ttc ggt ggt tgg aag aac tcc ctc ttc ggt gac gcc aag gca tat Ser Phe Gly Gly Trp Lys Asn Ser Leu Phe Gly Asp Ala Lys Ala Tyr 460 465 ggc act caa ggt ttt gat ttc ttc acc agg gaa aag gcg atc acc agc 1555 Gly Thr Gln Gly Phe Asp Phe Phe Thr Arg Glu Lys Ala Ile Thr Ser

470 475 480 485

cgt tgg ctc gac cca gca acc cac ggt ggc att aac ctc ggt ttc cca 1603

Arg Trp Leu Asp Pro Ala Thr His Gly Gly Ile Asn Leu Gly Phe Pro
490 495 500

cag aac gat taattgaagg agagcacagg act 1635 Gln Asn Asp

<210> 136

<211> 504

<212> PRT

<213> Corynebacterium glutamicum

<400> 136

Met Ser Glu Pro Gln Thr Ile Ser His Trp Ile Asp Gly Ala Ile Ser 1 5 10 15

Pro Ser Thr Ser Gly Lys Thr Ala Pro Val Tyr Asn Pro Ala Thr Gly 20 25 30

Gln Val Thr Ala Asn Val Ala Leu Ala Ser Gln Glu Glu Ile Asp Ala 35 40 45

Thr Ile Ala Ser Ala Thr Lys Ala Ala Lys Thr Trp Gly Asn Leu Ser 50 55 60

Ile Ala Lys Arg Gln Ala Val Leu Phe Asn Phe Arg Glu Leu Leu Asn 65 70 75 80

Ala Arg Lys Gly Glu Leu Ala Glu Ile Ile Thr Ala Glu His Gly Lys 85 90 95

Val Leu Ser Asp Ala Met Gly Glu Ile Leu Arg Gly Gln Glu Val Val
100 105 110

Glu Leu Ala Thr Gly Phe Pro His Leu Leu Lys Gly Ala Phe Asn Glu 115 120 125

Asn Val Ser Thr Gly Ile Asp Val Tyr Ser Leu Lys Gln Pro Leu Gly 130 135 140

Val Val Gly Ile Ile Ser Pro Phe Asn Phe Pro Ala Met Val Pro Met 145 150 155 160

Trp Phe Phe Pro Ile Ala Ile Ala Ala Gly Asn Ala Val Ile Leu Lys
165 170 175

Pro Ser Glu Lys Asp Pro Ser Ala Ala Leu Trp Met Ala Gln Ile Trp
180 185 190

Lys Glu Ala Gly Leu Pro Asp Gly Val Phe Asn Val Leu Gln Gly Asp 195 200 205

Lys Leu Ala Val Asp Gly Leu Leu Asn Ser Pro Asp Val Ser Ala Ile 210 215 220

Ser Phe Val Gly Ser Thr Pro Ile Ala Lys Tyr Ile Tyr Glu Thr Ser 230 Ala Lys Asn Gly Lys Arg Val Gln Ala Leu Gly Gly Ala Lys Asn His 250 Met Leu Val Leu Pro Asp Ala Asp Leu Asp Leu Val Ala Asp Gln Ala 265 Ile Asn Ala Gly Tyr Gly Ala Ala Gly Glu Arg Cys Met Ala Val Ser 280 Val Val Leu Ala Ile Glu Ser Val Ala Asp Glu Leu Ile Glu Lys Ile 295 Lys Glu Arg Ile Asp Thr Leu Arg Ile Gly Asn Gly Ala Gly Asp Glu 310 315 Gln Gly Glu Pro His Leu Gly Pro Leu Ile Thr Asp Val His Arg Asp 330 Lys Val Ala Ser Tyr Val Asp Ile Ala Glu Ala Asp Gly Ala Lys Ile 345 Ile Val Asp Gly Arg Asn Cys Ala Val Asp Gly His Glu Glu Gly Phe 360 Phe Phe Gly Pro Thr Leu Ile Asp Asp Ile Pro Leu Thr Phe Arg Ala 375 Tyr Thr Glu Glu Ile Phe Gly Pro Val Leu Ser Val Val Arg Val Ala 390 395 Ser Phe Asp Glu Ala Ile Glu Leu Ile Asn Ser Gly Glu Phe Gly Asn 410 Gly Thr Ala Ile Phe Thr Asn Asp Gly Gly Ala Ala Arg Arg Phe Gln 420 425 His Glu Ile Glu Val Gly Met Ile Gly Ile Asn Val Pro Ile Pro Val 440

Pro Val Ala Tyr His Ser Phe Gly Gly Trp Lys Asn Ser Leu Phe Gly 450 455 460

Asp Ala Lys Ala Tyr Gly Thr Gln Gly Phe Asp Phe Phe Thr Arg Glu

Lys Ala Ile Thr Ser Arg Trp Leu Asp Pro Ala Thr His Gly Gly Ile
485 490 495

475

Asn Leu Gly Phe Pro Gln Asn Asp 500

470

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<212> DNA

<213> Corynebacterium glutamicum

<220>

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		35					40					45				
Ile	Thr 50	Asn	Gly	Ala	Arg	Leu 55	Glu	Thr	Tyr	Val	Ile 60	Val	Gly	Asp	Ala	
Gly 65	Thr	Gly	Asn	Ile	Cys 70	Ile	Asn	Gly	Ala	Ala 75	Ala	His	Leu	Ile	Asn 80	
Pro	Gly	Asp	Leu	Val 85	Ile	Ile	Met	Ser	Tyr 90	Leu	Gln	Ala	Thr	Asp 95	Ala	
Glu	Ala	Lys	Ala 100	Tyr	Glu	Pro	Lys	Ile 105	Val	His	Val	Asp	Ala 110	Asp	Asn	
Arg	Ile	Val 115	Ala	Leu	Gly	Asn	Asp 120	Leu	Ala	Glu	Ala	Leu 125	Pro	Gly	Ser	
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gat	gtc										1				5	
Asp	Val	atc Ile	acc Thr	gcc Ala 10	caa Gln	caa Gln	cga Arg	acc Thr	gcc Ala 15	cct Pro	cat	gtt Val	cga Arg	cga Arg 20	5 acg Thr	163
cca	Val ctt Leu	Ile ttc	Thr	Ala 10 gca	Gln gac	Gln	Arg atc	Thr gac	Ala 15 ggc	Pro	cat His	Val atc	Arg tgg	Arg 20 atc Ile	5 acg Thr	163 211
cca Pro	ctt	Ile ttc Phe	Thr  gaa Glu 25 ctc	Ala 10 gca Ala caa	Gln gac Asp	Gln ccc Pro	atc Ile	Thr gac Asp 30	Ala 15 ggc Gly	Pro aca Thr	cat His caa Gln	Val atc Ile	tgg Trp 35	Arg 20 atc Ile gca	5 acg Thr aaa Lys	
cca Pro gca Ala	ctt Leu	ttc Phe ttc Phe	Thr  gaa Glu 25 ctc Leu	Ala 10 gca Ala caa Gln	gac Asp aag Lys	Gln ccc Pro tgc Cys	atc Ile ggc Gly 45 gaa Glu	gac Asp 30 gtg Val	Ala 15 ggc Gly ttc Phe	Pro aca Thr aaa Lys	cat His caa Gln acg Thr	atc Ile cgt Arg 50	tgg Trp 35 gga Gly	atc Ile gca Ala	acg Thr aaa Lys ttc Phe	211
cca Pro gca Ala aac Asn	ctt Leu gag Glu cgc Arg 55	ttc Phe ttc Phe 40 cag Gln	Thr gaa Glu 25 ctc Leu crc	Ala 10 gca Ala caa Gln gca Ala	gac Asp aag Lys gct Ala	ccc Pro tgc Cys tcg Ser 60 ggc	atc Ile ggc Gly 45 gaa Glu	gac Asp 30 gtg Val aac Asn	Ala 15 ggc Gly ttc Phe gga Gly	Pro aca Thr aaa Lys cta Leu	cat His caa Gln acg Thr ctc Leu 65	atc Ile cgt Arg 50 gac Asp	tgg Trp 35 gga Gly cca Pro	atc Ile gca Ala acg Thr	acg Thr aaa Lys ttc Phe	211

_			_		-	-	cgc Arg		-							451
					_		gcg Ala 125	_	-		-	-	-			499
	-	-	-			-	ctg Leu		-		-		_	_		547
							gtc Val									595
		-	_	-			gtg Val	-	_	-						643
	_			_	-	_	gta Val	-	_		-					691
_	_	_					cca Pro 205		_						_	739
			~		-		gtt Val					-	-		_	787
	_	_				_	gaa Glu	_		-		-		-		835
							gac Asp									883
_				_			cgc			_						931
ñгg	1113	ьеч	265	ASD	ASII	ıyı	AIG	270	Pro	Ala	Glu	His	275	Ala	Ala	
gca	gca	ctc	gcc	tct	ctt	acc	agt	gga	gca	tac	aaa	cct	gca	gca	gat	979
Ala	Ala	Leu 280	Ala	Ser	Leu	Thr	Ser 285		Ala	Tyr	Lys	Pro 290	Ala	Ala	Asp	
gaa 1023		gtg	gca	gtc	att	gtg	tgc	gga	gcg	aac	act	gac	ctc	aca	aca	
		Val	Ala	Val	Ile	Val 300	Cys	Gly	Ala	Asn	Thr 305	Asp	Leu	Thr	Thr	
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310																

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<211> 310

<212> PRT

<213> Corynebacterium glutamicum

<400> 140

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His Val Arg Arg Thr Pro Leu Phe Glu Ala Asp Pro Ile Asp Gly Thr 20 25 30

Gln Ile Trp Ile Lys Ala Glu Phe Leu Gln Lys Cys Gly Val Phe Lys  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Thr Arg Gly Ala Phe Asn Arg Gln Leu Ala Ala Ser Glu Asn Gly Leu 50 55 60

Leu Asp Pro Thr Val Gly Ile Val Ala Ala Ser Gly Gly Asn Ala Gly 65 70 75 80

Leu Ala Asn Ala Phe Ala Ala Ala Ser Leu Ser Val Pro Ala Thr Val 85 90 95

Leu Val Pro Glu Thr Ala Pro Gln Val Lys Val Asp Arg Leu Lys Gln 100 105 110

Tyr Gly Ala Thr Val Gln Gln Ile Gly Ser Glu Tyr Ala Glu Ala Phe 115 120 125

Glu Ala Ala Gln Thr Phe Glu Ser Glu Thr Gly Ala Leu Phe Cys His 130 135 140

Ala Tyr Asp Gln Pro Asp Ile Ala Ala Gly Ala Gly Val Ile Gly Leu 145 150 155 160

Glu Ile Val Glu Asp Leu Pro Asp Val Asp Thr Ile Val Val Ala Val
165 170 175

Gly Gly Gly Leu Tyr Ala Gly Ile Ala Ala Val Val Ala Ala His 180 185 190

Asp Ile Lys Val Val Ala Val Glu Pro Ser Lys Ile Pro Thr Leu His 195 200 205

Asn Ser Leu Ile Ala Gly Gln Pro Val Asp Val Asn Val Ser Gly Ile 210 215 220

Ala Ala Asp Ser Leu Gly Ala Arg Gln Ile Gly Arg Glu Ala Phe Asp 225 230 235 240

Ile Ala Thr Ala His Pro Pro Ile Gly Val Leu Val Asp Asp Glu Ala 245 250 255

Ile Ile Ala Ala Arg Arg His Leu Trp Asp Asn Tyr Arg Ile Pro Ala 260 265 270

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Lys Pro Ala Ala Asp Glu Lys Val Ala Val Ile Val Cys Gly Ala Asn 290 295 300

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toc ca Ser G	aa ln	tta Leu	ctc Leu	gca Ala 170	tat Tyr	ggt Gly	cgc Arg	gat Asp	ttt Phe 175	gcg Ala	gag Glu	gtc Val	atg Met	aag Lys 180	gat Asp	643
aat g Asn G	ag lu	cgc Arg	tta Leu 185	atc Ile	cac His	Gly ggg	gat Asp	ctt Leu 190	ggc Gly	aca Thr	gtg Vaî	gat Asp	gcc Ala 195	cat His	ttg Leu	691
gat c Asp A	ga .rg	gtg Val 200	tgg Trp	cag Gln	att Ile	atg Met	cag Gln 205	gag Glu	tgc Cys	gtg Val	gca Ala	caa Gln 210	ggc Gly	atc Ile	gca Ala	739
acg c Thr P 2	cg ro	Gly	att Ile	tta Leu	ccg Pro	ggt Gly 220	ggg Gly	ttg Leu	aat Asn	gtg Val	caa Gln 225	cgt Arg	cgg Arg	gcg Ala	ccg Pro	787
cag g Gln V 230	ta al	cac His	gcg Ala	ctg Leu	att Ile 235	agc Ser	aac Asn	ggg	gat Asp	acg Thr 240	tgt Cys	gag Glu	ctg Leu	ggt Gly	gct Ala 245	835
gat c Asp L	tt Leu	gat Asp	gct Ala	gtg Val 250	gag Glu	tgg Trp	gtg Val	aat Asn	ctg Leu 255	tac Tyr	gcc Ala	ttg Leu	gcg Ala	gtg Val 260	aat Asn	883
gaa g Glu G	gaa Glu	aac Asn	gcc Ala 265	gct Ala	ggt Gly	ggt Gly	cgt Arg	gtg Val 270	gtt Val	act Thr	gct Ala	ccg Pro	act Thr 275	aat Asn	ggt Gly	931
gct g Ala A	gcg Ala	ggg Gly 280	att Ile	att Ile	ccg Pro	gcg Ala	gtg Val 285	atg Met	cac His	tat Tyr	gcg Ala	cgg Arg 290	gat Asp	ttt Phe	ttg Leu	979
aca g	ggt	ttt	ggg	gcg	gag	cag	gcg	cgg	acg	ttt	ttg	tat	acc	gcg	ggt	
1027 Thr C	Gly 295	Phe	Gly	Ala	Glu	Gln 300	Ala	Arg	Thr	Phe	Leu 305	Tyr	Thr	Ala	Gly	
gcg 9	gtg	ggc	atc	atc	att	aag	gaa	aat	gcc	tcg	atc	tct	ggc	gcg	gag	
Ala \ 310	Val	Gly	Ile	Ile	Ile 315	Lys	Glu	Asn	Ala	Ser 320		Ser	Gly	Ala	Glu 325	
gtg (	ggg	tgt	cag	ggt	gag	gtt	ggt	tca	gcg	tcc	gcg	atg	gcg	gct	gcc	
Val (	Gly	Cys	Gln	Gly 330		Val	Gly	Ser	Ala 335		Ala	Met	Ala	Ala 340		
ggg 1		tgt	gca	gtc	tta	ggt	ggt	tct	ccg	caa	cag	gtg	gaa	aac	gcc	
Gly 1	Leu	Суѕ	Ala 345		Leu	Gly	Gly	Ser 350		Gln	Gln	Val	Glu 355	Asn	Ala	
gcg (		att	gcg	ttg	gag	cac	aat	ttg	gga	ttg	, acg	tgc	gat	ccg	gtg	
Ala (		Ile 360		Leu	Glu	His	Asn 365		Gly	Leu	Thr	Cys 370		Pro	Val	
ggc		tta	gtg	r caç	att	ccg	tgt	att	gaa	. cgc	aac	gct	att	gct	gcc	
1267 Gly	Gly	Leu	. Val	. Glr	ı Ile	Pro	Cys	Ile	Glu	Arç	g Asr	n Ala	ıl∈	e Ala	Ala	

375 380 385

atg aag too atc aat gog goa agg ott goo ogg att ggt gat gg<br/>c aac 1315

Met Lys Ser Ile Asn Ala Ala Arg Leu Ala Arg Ile Gly Asp Gly Asn 390 395 400 405

aat cgc gtg agt ttg gat gtg gtg gtc acg atg gct gcc acc ggc 1363

Asn Arg Val Ser Leu Asp Asp Val Val Val Thr Met Ala Ala Thr Gly 410 415 420

cgg gac atg ctg acc aaa tat aag gaa acg tcc ctt ggt ggt ttg gca 1411

Arg Asp Met Leu Thr Lys Tyr Lys Glu Thr Ser Leu Gly Gly Leu Ala 425 430 435

acc acc ttg ggc ttc ccg gtg tcg atg acg gag tgt tagcggtacg 1457

Thr Thr Leu Gly Phe Pro Val Ser Met Thr Glu Cys
440

gctttaacac ggc 1470

<210> 142

<211> 449

<212> PRT

<213> Corynebacterium glutamicum

<400> 142

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Ser Ser His Thr Val Gly Pro Met Arg Ala Ala Leu Thr Tyr Ile Ser 20 25 30

Glu Phe Pro Ser Ser His Val Asp Ile Thr Leu His Gly Ser Leu Ala 35 40 45

Ala Thr Gly Lys Gly His Cys Thr Asp Arg Ala Val Leu Leu Gly Leu

Val Gly Trp Glu Pro Thr Ile Val Pro Ile Asp Ala Ala Pro Ser Pro

Gly Ala Pro Ile Pro Ala Lys Gly Ser Val Asn Gly Pro Lys Gly Thr 85 90 95

Val Ser Tyr Ser Leu Thr Phe Asp Pro His Pro Leu Pro Glu His Pro
100 105 110

Asn Ala Val Thr Phe Lys Gly Ser Thr Thr Arg Thr Tyr Leu Ser Val 115 120 125

Gly Gly Phe Ile Met Thr Leu Glu Asp Phe Arg Lys Leu Asp Asp 130 135 140

Ile Gly Ser Gly Val Ser Thr Ile His Pro Glu Ala Glu Val Pro Cys 145 150 155 160 Pro Phe Gln Lys Ser Ser Gln Leu Leu Ala Tyr Gly Arg Asp Phe Ala 165 170 175

- Glu Val Met Lys Asp Asn Glu Arg Leu Ile His Gly Asp Leu Gly Thr 180 185 190
- Val Asp Ala His Leu Asp Arg Val Trp Gln Ile Met Gln Glu Cys Val 195 200 205
- Ala Gln Gly Ile Ala Thr Pro Gly Ile Leu Pro Gly Gly Leu Asn Val 210 215 220
- Gln Arg Arg Ala Pro Gln Val His Ala Leu Ile Ser Asn Gly Asp Thr 225 230 235 240
- Cys Glu Leu Gly Ala Asp Leu Asp Ala Val Glu Trp Val Asn Leu Tyr 245 250 255
- Ala Leu Ala Val Asn Glu Glu Asn Ala Ala Gly Gly Arg Val Val Thr 260 265 270
- Ala Pro Thr Asn Gly Ala Ala Gly Ile Ile Pro Ala Val Met His Tyr 275 280 285
- Ala Arg Asp Phe Leu Thr Gly Phe Gly Ala Glu Gln Ala Arg Thr Phe 290 295 300
- Leu Tyr Thr Ala Gly Ala Val Gly Ile Ile Ile Lys Glu Asn Ala Ser 305 310 315 320
- Ile Ser Gly Ala Glu Val Gly Cys Gln Gly Glu Val Gly Ser Ala Ser
- Ala Met Ala Ala Gly Leu Cys Ala Val Leu Gly Gly Ser Pro Gln 340 345 350
- Gln Val Glu Asn Ala Ala Glu Ile Ala Leu Glu His Asn Leu Gly Leu 355 360 365
- Thr Cys Asp Pro Val Gly Gly Leu Val Gln Ile Pro Cys Ile Glu Arg 370 375 380
- Asn Ala Ile Ala Ala Met Lys Ser Ile Asn Ala Ala Arg Leu Ala Arg 385 390 395 400
- Ile Gly Asp Gly Asn Asn Arg Val Ser Leu Asp Asp Val Val Thr 405 410 415
- Met Ala Ala Thr Gly Arg Asp Met Leu Thr Lys Tyr Lys Glu Thr Ser 420 425 430
- Leu Gly Gly Leu Ala Thr Thr Leu Gly Phe Pro Val Ser Met Thr Glu 435 440 445

Cys

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185		190		195
gct gcg gaa gtt Ala Ala Glu Val 200	Gly Ala Lys	ctg tgg gtc Leu Trp Val 205	gat atg gct Asp Met Ala 210	cac ttc gct 739 His Phe Ala
ggt ctt gtt gct Gly Leu Val Ala 215	gct ggt ttg Ala Gly Leu 220	cac cca agc His Pro Ser	cca gtt cct Pro Val Pro 225	tac tct gat 787 Tyr Ser Asp
gtt gtt tct tcc Val Val Ser Ser 230	act gtc cac Thr Val His 235	aag act ttg Lys Thr Leu	ggt gga cct Gly Gly Pro 240	cgt tcc ggc 835 Arg Ser Gly 245
atc att ctg gct Ile Ile Leu Ala	aag cag gag Lys Gln Glu 250	tac gcg aag Tyr Ala Lys 255	aag ctg aac Lys Leu Asn	tct tcc gta 883 Ser Ser Val 260
ttc cca ggt cag Phe Pro Gly Gln 265	cag ggt ggt Gln Gly Gly	cct ttg atg Pro Leu Met 270	cac gca gtt His Ala Val	gct gcg aag 931 Ala Ala Lys 275
gct act tct ttg Ala Thr Ser Leu 280	aag att gct Lys Ile Ala	ggc act gag Gly Thr Glu 285	cag ttc cgt Gln Phe Arg 290	gac cgt cag 979 Asp Arg Gln
gct cgc acg ttg	gag ggt gct	cgc att ctt	gct gag cgt	ctg act gct
Ala Arg Thr Leu 295	Glu Gly Ala 300	Arg Ile Leu	Ala Glu Arg 305	Leu Thr Ala
tct gat gcg aag	gcc gct ggc	gtg gat gtc	ttg acc ggt	ggc act gat
1075 Ser Asp Ala Lys 310	Ala Ala Gly 315	Val Asp Val	Leu Thr Gly 320	Gly Thr Asp 325
gtg cac ttg gtt 1123	ttg gct gat	ctg cgt aac	tcc cag atg	gat ggc cag
Val His Leu Val	Leu Ala Asp 330	Leu Arg Asr 335		Asp Gly Gln 340
cag gcg gaa gat	ctg ctg cac	gag gtt ggt	atc act gtg	aac cgt aac
1171 Gln Ala Glu Asp 345		Glu Val Gly 350	lle Thr Val	Asn Arg Asn 355
gcg gtt cct ttc 1219	gat cct cgt	cca cca ato	gtt act tct	ggt ctg cgt
Ala Val Pro Phe	Asp Pro Arg	Pro Pro Met 365	Val Thr Ser 370	Gly Leu Arg
att ggt act cct	gcg ctg gct	acc cgt ggt	ttc gat att	cct gca ttc
1267 Ile Gly Thr Pro 375	Ala Leu Ala 380	Thr Arg Gly	Phe Asp Ile 385	Pro Ala Phe
act gag gtt gca	gac atc att	ggt act gc	ttg gct aat	ggt aag tcc
1315 Thr Glu Val Ala 390	A Asp Ile Ile 395	Gly Thr Ala	a Leu Ala Asn 400	Gly Lys Ser 405

gca gac att gag tet etg egt gge egt gta gea aag ett get gea gat 1363

Ala Asp Ile Glu Ser Leu Arg Gly Arg Val Ala Lys Leu Ala Ala Asp 410 415 420

tac cca ctg tat gag ggc ttg gaa gac tgg acc atc gtc taagtttttc 1412

Tyr Pro Leu Tyr Glu Gly Leu Glu Asp Trp Thr Ile Val 425

tttgagtttt cat 1425

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<211> 434

<212> PRT

<213> Corynebacterium glutamicum

<400> 144

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Gln Arg Asp Thr Leu Glu Met Ile Ala Ser Glu Asn Phe Val Pro Arg 35 40 45

Ser Val Leu Gln Ala Gln Gly Ser Val Leu Thr Asn Lys Tyr Ala Glu 50 55 60

Gly Tyr Pro Gly Arg Arg Tyr Tyr Gly Gly Cys Glu Gln Val Asp Ile 65 70 75 80

Ile Glu Asp Leu Ala Arg Asp Arg Ala Lys Ala Leu Phe Gly Ala Glu 85 90 95

Phe Ala Asn Val Gln Pro His Ser Gly Ala Gln Ala Asn Ala Ala Val 100 105 110

Leu Met Thr Leu Ala Glu Pro Gly Asp Lys Ile Met Gly Leu Ser Leu

Ala His Gly Gly His Leu Thr His Gly Met Lys Leu Asn Phe Ser Gly 130 135 140

Val Asp Met Asp Gln Val Arg Glu Ile Ala Leu Lys Glu Gln Pro Lys 165 170 175

Val Ile Ile Ala Gly Trp Ser Ala Tyr Pro Arg His Leu Asp Phe Glu 180 185 190

Ala Phe Gln Ser Ile Ala Ala Glu Val Gly Ala Lys Leu Trp Val Asp 195 200 205

Met Ala His Phe Ala Gly Leu Val Ala Ala Gly Leu His Pro Ser Pro 210 215 220

Val Pro Tyr Ser Asp Val Val Ser Ser Thr Val His Lys Thr Leu Gly 230 Gly Pro Arg Ser Gly Ile Ile Leu Ala Lys Gln Glu Tyr Ala Lys Lys Leu Asn Ser Ser Val Phe Pro Gly Gln Gln Gly Gly Pro Leu Met His Ala Val Ala Ala Lys Ala Thr Ser Leu Lys Ile Ala Gly Thr Glu Gln 280 Phe Arg Asp Arg Gln Ala Arg Thr Leu Glu Gly Ala Arg Ile Leu Ala Glu Arg Leu Thr Ala Ser Asp Ala Lys Ala Ala Gly Val Asp Val Leu 315 310 Thr Gly Gly Thr Asp Val His Leu Val Leu Ala Asp Leu Arg Asn Ser 330 325 Gln Met Asp Gly Gln Gln Ala Glu Asp Leu Leu His Glu Val Gly Ile 345 340 Thr Val Asn Arg Asn Ala Val Pro Phe Asp Pro Arg Pro Pro Met Val 360 Thr Ser Gly Leu Arg Ile Gly Thr Pro Ala Leu Ala Thr Arg Gly Phe 375 Asp Ile Pro Ala Phe Thr Glu Val Ala Asp Ile Ile Gly Thr Ala Leu 390 Ala Asn Gly Lys Ser Ala Asp Ile Glu Ser Leu Arg Gly Arg Val Ala 410 405 Lys Leu Ala Ala Asp Tyr Pro Leu Tyr Glu Gly Leu Glu Asp Trp Thr 425 Ile Val <210> 145 <211> 401 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (1)..(378) <223> RXA01821 <400> 145 cga aac agc caa ggc aaa tgg tgc cca agt acg cga tca cca aaa aat Arg Asn Ser Gln Gly Lys Trp Cys Pro Ser Thr Arg Ser Pro Lys Asn 10 1 -5 acc agc atc gaa gac aac ggc gat cac gta gtc atc caa gca ggc gaa Thr Ser Ile Glu Asp Asn Gly Asp His Val Val Ile Gln Ala Gly Glu

			20					25					30			
-				-	gac Asp	_										144
					tcc Ser											192
					ctg Leu 70											240
					atc Ile											288
					Gly											336
		_			agc Ser				_							378
tga	tgtc	ctg a	atcc	ggt	tc c	gg										401
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Thr	Ser	Ile	Glu 20	Asp	Asn	Gly	Asp	His 25	Val	Val	Ile	Gln	Ala 30	Gly	Glu	
Glu	Thr	Thr	Ile	Val	Asp	Arg	Val	Ile	Val	Thr	Thr	Gly 45	Ser	Trp	Thr	
Ser	Glu 50	Leu	Val	Pro	Ser	Ile 55	Ala	Pro	Leu	Leu	Glu 60	Val	Arg	Arg	Leu	
Val 65	Leu	Thr	Trp	Phe	Leu 70	Pro	Asn	Asn	Pro	Val 75	Asp	Phe	Gln	Pro	Glu 80	
Asn	Leu	Pro	Cys	Phe 85	Ile	Arg	Asp	Arg	Asp 90	Gly	Phe	His	Val	Phe 95	Gly	
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atc Ile	ggc Gly	ctt Leu	gga Gly	tca Ser 10	acc Thr	ggc Gly	tcc Ser	atg Met	gca Ala 15	ctg Leu	tgg Trp	cac His	tta Leu	agt Ser 20	aac Asn	163
atc Ile	cca Pro	ggt Gly	gta Val 25	gag Glu	gcc Ala	atc Ile	ggc Gly	ttt Phe 30	gaa Glu	caa Gln	ttc Phe	ggc Gly	atc Ile 35	tcc Ser	cat His	211
ggc Gly	tac Tyr	ggc Gly 40	gca Ala	ttc Phe	aca Thr	ggg Gly	gag Glu 45	tcc Ser	cga Arg	ctg Leu	ttt Phe	cgc Arg 50	atg Met	gcc Ala	tac Tyr	259
cac His	gaa Glu 55	ggc Gly	agc Ser	acc Thr	tac Tyr	gtt Val 60	ccg Pro	ttg Leu	ctc Leu	aaa Lys	cgc Arg 65	gca Ala	cga Arg	gca Ala	cta Leu	307
tgg Trp 70	tca Ser	tca Ser	ctg Leu	agc Ser	gag Glu 75	att Ile	tcc Ser	gga Gly	cgc Arg	gaa Glu 80	ctc Leu	ttc Phe	cac His	aac Asn	ttc Phe 85	355
ggt Gly	gtc Val	tta Leu	agc Ser	acc Thr 90	ggc Gly	aag Lys	gaa Glu	gac Asp	gaa Glu 95	gca Ala	ccc Pro	ttc Phe	caa Gln	cgc Arg 100	ctg Leu	403
gtg Val	gaa Glu	tca Ser	gtg Val 105	gaa Glu	cgt Arg	tat Tyr	gag Glu	ctg Leu 110	cca Pro	cat His	gaa Glu	cga Arg	ctt Leu 115	Thr	gcc Ala	451
gcg Ala	cag Gln	atg Met 120	cgc Arg	agc Ser	gtt Val	acc Thr	cag Gln 125	Val	tag	actt	ccg					488
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Trp	His	: Let	Ser 20		Ile	Pro	Gly	/ Val 25	Glu	ı Ala	a Ile	e Gly	Phe	e Glu	Gln	

Phe Gly Ile Ser His Gly Tyr Gly Ala Phe Thr Gly Glu Ser Arg Leu Phe Arg Met Ala Tyr His Glu Gly Ser Thr Tyr Val Pro Leu Leu Lys 50 55 Arg Ala Arg Ala Leu Trp Ser Ser Leu Ser Glu Ile Ser Gly Arg Glu 7.0 Leu Phe His Asn Phe Gly Val Leu Ser Thr Gly Lys Glu Asp Glu Ala Pro Phe Gln Arg Leu Val Glu Ser Val Glu Arg Tyr Glu Leu Pro His 105 Glu Arg Leu Thr Ala Ala Gln Met Arg Ser Val Thr Gln Val 120 <210> 149 <211> 460 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(460) <223> FRXA02263 <400> 149 cctgggcaac ccaagtgtat gaaaacgccc tggaaaaagg cgtcggcacc acattgaacc 60 tgtgggaatc accegcactg gettgagaga agaaacaaca atg aaa att geg gta Met Lys Ile Ala Val 1 atc ggc ctt gga tca acc ggc tcc atg gca ctg tgg cac tta agt aac Ile Gly Leu Gly Ser Thr Gly Ser Met Ala Leu Trp His Leu Ser Asn atc cca ggt gta gag gcc atc ggc ttt gaa caa ttc ggc atc tcc cat Ile Pro Gly Val Glu Ala Ile Gly Phe Glu Gln Phe Gly Ile Ser His gge tae gge gea tte aca ggg gag tee ega etg ttt ege atg gee tae 259 Gly Tyr Gly Ala Phe Thr Gly Glu Ser Arg Leu Phe Arg Met Ala Tyr 40 50 cac gaa ggc agc acc tac gtt ccg ttg ctc aaa cgc gca cga gca cta 307 His Glu Gly Ser Thr Tyr Val Pro Leu Leu Lys Arg Ala Arg Ala Leu 55 tgg tca tca ctg age gag att tcc gga cgc gaa ctc ttc cac aac ttc Trp Ser Ser Leu Ser Glu Ile Ser Gly Arg Glu Leu Phe His Asn Phe ggt gtc tta agc acc ggc aag gaa gac gaa gca ccc ttc caa cgc ctg 403 Gly Val Leu Ser Thr Gly Lys Glu Asp Glu Ala Pro Phe Gln Arg Leu 90 95 gtg gaa tca gtg gaa cgt tat gag ctg cca cat gaa cga ctt acc gcc 451

Val Glu Ser Val Glu Arg Tyr Glu Leu Pro His Glu Arg Leu Thr Ala 110 460 gcg cag atg Ala Gln Met 120 <210> 150 <211> 120 <212> PRT <213> Corynebacterium glutamicum Met Lys Ile Ala Val Ile Gly Leu Gly Ser Thr Gly Ser Met Ala Leu Trp His Leu Ser Asn Ile Pro Gly Val Glu Ala Ile Gly Phe Glu Gln Phe Gly Ile Ser His Gly Tyr Gly Ala Phe Thr Gly Glu Ser Arg Leu Phe Arg Met Ala Tyr His Glu Gly Ser Thr Tyr Val Pro Leu Leu Lys Arg Ala Arg Ala Leu Trp Ser Ser Leu Ser Glu Ile Ser Gly Arg Glu Leu Phe His Asn Phe Gly Val Leu Ser Thr Gly Lys Glu Asp Glu Ala Pro Phe Gln Arg Leu Val Glu Ser Val Glu Arg Tyr Glu Leu Pro His 105 100 Glu Arg Leu Thr Ala Ala Gln Met 115 <210> 151 <211> 1251 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1228) <223> RXA02176 <400> 151 gggtgctagg aactgacage ttcagggtta tagttgttgq gtcagategt taacgatece 60 tggccctttt acttccaage gcagaaagtt gcccgaagac atg ace gac tte ccc Met Thr Asp Phe Pro acc ctg ccc tct gag ttc atc cct ggc gac ggc cgt ttc ggc tgc gga 163 Thr Leu Pro Ser Glu Phe Ile Pro Gly Asp Gly Arg Phe Gly Cys Gly 10 cct tcc aag gtt cga cca gaa cag att cag gct att gtc gac gga tcc

110	Ser	Lys	Val 25	Arg	Pro	Glu	Gln	Ile 30	Gln	Ala	Ile	Val	Asp 35	Gly	Ser	
-				ggt Gly				-	_	_					_	259
				cgc Arg												307
				atc Ile												355
				gga Gly 90												403
				tcc Ser	_		_	_	-		-		_			451
				gag Glu												499
				gaa Glu												547
				gcc Ala	_	-		-		_	ccc Pro					595
Thr 150 ggc	Ser	Thr ctg	Gly	-	Met 155 att	Val gac	Pro gca	Val	Leu	Arg 160 ggc	Pro gct	Glu ggt	Gly	Ser ctg	Glu 165 cca	<ul><li>595</li><li>643</li></ul>
Thr 150 ggc Gly	ser tcc ser	Thr ctg Leu	Gly gtt Val	Ala gcc Ala	Met 155 att Ile	Val gac Asp gat Asp	Pro gca Ala gtt	Val acc Thr tac	tcc Ser 175 tac Tyr	Arg 160 ggc Gly	Pro gct Ala tcc	Glu ggt Gly cca Pro	Gly gga Gly cag	ctg Leu 180 aag Lys	Glu 165 cca Pro	
Thr 150 ggc Gly gta Val	Ser tcc Ser gac Asp	Thr ctg Leu atc Ile	Gly gtt Val aag Lys 185	Ala gcc Ala 170 aac	Met 155 att Ile tcc Ser	Val gac Asp gat Asp	gca Ala gtt Val	val acc Thr tac Tyr 190	tcc Ser 175 tac Tyr	Arg 160 ggc Gly ttc Phe	Pro gct Ala tcc Ser	ggt Gly cca Pro	gga Gly cag Gln 195	ctg Leu 180 aag Lys	Glu 165 cca Pro tgc Cys	643
Thr 150 ggc Gly gta Val ttc Phe	tcc Ser gac Asp gca Ala	Thr ctg Leu atc Ile tcc Ser 200 cgc	gtt Val aag Lys 185 gac Asp	Ala gcc Ala 170 aac Asn	Met 155 att Ile tcc Ser ggc Gly	yal gac Asp gat Asp ctg Leu	gca Ala gtt Val tgg Trp 205	acc Thr tac Tyr 190 ctt Leu	tcc Ser 175 tac Tyr gca Ala	Arg 160 ggc Gly ttc Phe gcg Ala	gct Ala tcc Ser atg Met cgc	ggt Gly cca Pro agc Ser 210	gga Gly cag Gln 195 cca Pro	ctg Leu 180 aag Lys gca Ala	Glu 165 cca Pro tgc Cys	643
Thr 150  ggc Gly  gta Val  ttc Phe  ctc Leu	tcc Ser gac Asp gca Ala gag Glu 215	Thr  ctg Leu  atc Ile  tcc Ser 200  cgc Arg	Gly gtt Val aag Lys 185 gac Asp atc Ile	gcc Ala 170 aac Asn ggt Gly	Met 155 att Ile tcc Ser ggc Gly aag Lys	yal gac Asp gat Asp ctg Leu atc Ile 220 gca	gca Ala gtt Val tgg Trp 205 aac Asn	Val acc Thr tac Tyr 190 ctt Leu gct Ala	tcc Ser 175 tac Tyr gca Ala tcc Ser	Arg 160 ggc Gly ttc Phe gcg Ala gat Asp	gct Ala tcc Ser atg Met cgc Arg 225 ctg	ggt Gly cca Pro agc Ser 210 ttc Phe	Gly gga Gly cag Gln 195 cca Pro atc Ile	ctg Leu 180 aag Lys gca Ala cct Pro	Glu 165 cca Pro tgc Cys gct Ala gag Glu	643 691 739
Thr 150  ggc Gly  gta Val  ttc Phe  ctc Leu  ttc Phe 230  tac	tcc Ser gac Asp gca Ala gag Glu 215 ctc Leu	Thr  ctg Leu  atc Ile  tcc Ser 200  cgc Arg  aac Asn	gtt Val aag Lys 185 gac Asp atc Ile ctg Leu	gcc Ala 170 aac Asn ggt Gly gag Glu cag	Met 155 att Ile tcc Ser ggc Gly aag Lys acc Thr 235	yal gac Asp gat Asp ctg Leu atc Ile 220 gca Ala gct	gca Ala gtt Val tgg Trp 205 aac Asn gtg Val	Val acc Thr tac Tyr 190 ctt Leu gct Ala gat Asp	tcc Ser 175 tac Tyr gca Ala tcc Ser aac Asn	Arg 160 ggc Gly ttc Phe gcg Ala gat Asp tcc Ser 240 atg	gct Ala tcc Ser atg Met cgc Arg 225 ctg Leu ctg	ggt Gly cca Pro agc Ser 210 ttc Phe aag Lys	gga Gly cag Gln 195 cca Pro atc Ile aac Asn	ctg Leu 180 aag Lys gca Ala cct Pro	Glu 165 cca Pro tgc Cys gct Ala gag Glu acc Thr 245	643 691 739 787

275 270 265 aca gca ago too too goo otg tao aac tgg got gag got ogo gag gag Thr Ala Ser Ser Ser Ala Leu Tyr Asn Trp Ala Glu Ala Arg Glu Glu 285 gca tee eea tae gtg gea gat gea get aag ege tee ete gtt gte gge 1027 Ala Ser Pro Tyr Val Ala Asp Ala Ala Lys Arg Ser Leu Val Val Gly 295 acc atc gac ttc gat gac tcc atc gac gca gca gtg atc gct aag ata Thr Ile Asp Phe Asp Asp Ser Ile Asp Ala Ala Val Ile Ala Lys Ile 315 ctg cgc gca aac ggc atc ctg gac acc gag cct tac cgc aag ctg gga Leu Arg Ala Asn Gly Ile Leu Asp Thr Glu Pro Tyr Arg Lys Leu Gly 330 cgc aac cag ctg cgc atc ggt atg ttc cca gcg atc gat tcc acc gat Arg Asn Gln Leu Arg Ile Gly Met Phe Pro Ala Ile Asp Ser Thr Asp 350 345 gtg gaa aag ctc acc gga gca atc gac ttc atc ctc gat ggc ggt ttt Val Glu Lys Leu Thr Gly Ala Ile Asp Phe Ile Leu Asp Gly Gly Phe 365 gca agg aag taataccccc actttgaaaa aca 1251 Ala Arg Lys 375 <210> 152 <211> 376 <212> PRT <213> Corynebacterium glutamicum <400> 152 Met Thr Asp Phe Pro Thr Leu Pro Ser Glu Phe Ile Pro Gly Asp Gly Arg Phe Gly Cys Gly Pro Ser Lys Val Arg Pro Glu Gln Ile Gln Ala 2.0 Ile Val Asp Gly Ser Ala Ser Val Ile Gly Thr Ser His Arg Gln Pro 40 Ala Val Lys Asn Val Val Gly Ser Ile Arg Glu Gly Leu Ser Asp Leu Phe Ser Leu Pro Glu Gly Tyr Glu Ile Ile Leu Ser Leu Gly Gly Ala Thr Ala Phe Trp Asp Ala Ala Thr Phe Gly Leu Ile Glu Lys Lys Ser

9.0

Gly His Leu Ser Phe Gly Glu Phe Ser Ser Lys Phe Ala Lys Ala Ser Lys Leu Ala Pro Trp Leu Asp Glu Pro Glu Ile Val Thr Ala Glu Thr Gly Asp Ser Pro Ala Pro Gln Ala Phe Glu Gly Ala Asp Val Ile Ala Trp Ala His Asn Glu Thr Ser Thr Gly Ala Met Val Pro Val Leu Arg 145 150 155 Pro Glu Gly Ser Glu Gly Ser Leu Val Ala Ile Asp Ala Thr Ser Gly 170 Ala Gly Gly Leu Pro Val Asp Ile Lys Asn Ser Asp Val Tyr Tyr Phe 180 185 190 Ser Pro Gln Lys Cys Phe Ala Ser Asp Gly Gly Leu Trp Leu Ala Ala 200 Met Ser Pro Ala Ala Leu Glu Arg Ile Glu Lys Ile Asn Ala Ser Asp 215 Arg Phe Ile Pro Glu Phe Leu Asn Leu Gln Thr Ala Val Asp Asn Ser 230 235 Leu Lys Asn Gln Thr Tyr Asn Thr Pro Ala Val Ala Thr Leu Leu Met 245 250 Leu Asp Asn Gln Val Lys Trp Met Asn Ser Asn Gly Gly Leu Asp Gly 265 Met Val Ala Arg Thr Thr Ala Ser Ser Ser Ala Leu Tyr Asn Trp Ala 280 Glu Ala Arg Glu Glu Ala Ser Pro Tyr Val Ala Asp Ala Ala Lys Arg Ser Leu Val Val Gly Thr Ile Asp Phe Asp Asp Ser Ile Asp Ala Ala 310 315 THE ALE LYS THE LEW AYE ALE ASH GIV THE LEW ASP THY GIW Pro 325 330 Tyr Arg Lys Leu Gly Arg Asn Gln Leu Arg Ile Gly Met Phe Pro Ala 345 350 Ile Asp Ser Thr Asp Val Glu Lys Leu Thr Gly Ala Ile Asp Phe Ile 360 365 Leu Asp Gly Gly Phe Ala Arg Lys 370 375 <210> 153 <211> 1422

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	200					205					210				
ttg atc Leu Ile 215			• •	_			_	_		-				-	787
gaa gct Glu Ala 230	-	-		-	-			-	-	_	_				835
gat ttc Asp Phe															883
gat gcg Asp Ala	_			_		-	_	_	-			_			931
ggt gcg Gly Ala	_		_			_	_								979
gct gtt 1027	gtt	tcc	ggt	ggt	ttc	atc	cag	gtg	ttg	gaa	ggt	ttg	gct	gag	
Ala Val 295	Val	Ser	Gly	Gly	Phe 300	Ile	Gln	Val	Leu	Glu 305	Gly	Leu	Ala	Glu	
gag ttg 1075	gag	ttg	gat	tat	gtc	cgc	gcc	aac	act	ttg	gaa	atc	gtt	gat	
Glu Leu 310	Glu	Leu	Asp	Туr 315	Val	Arg	Ala	Asn	Thr 320	Leu	Glu	Ile	Val	Asp 325	
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Gly Lys	Leu	Thr	Gly 330	Asn	Val	Thr	Gly	Lys 335	Ile	Val	Asp	Arg	Ala 340	Ala	
aag gct 1171	gag	ttc	ctc	cgt	gag	ttc	gct	gcg	gat	tct	ggc	ctg	aag	atg	
Lys Ala	Glu	Phe 345	Leu	Arg	Glu	Phe	Ala 350	Ala	Asp	Ser	Gly	Leu 355	Lys	Met	
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Tyr Gln	Thr 360	Val	Ala	Val	Gly	Asp 365	Gly	Ala	Asn	Asp	Ile 370	Asp	Met	Leu	
tcc gct 1267	gcg	ggt	ctg	ggt	gtt	gct	ttc	aac	gcg	aag	cct	gcg	ctg	aag	
Ser Ala 375	Ala	Gly	Leu	Gly	Val 380	Ala	Phe	Asn	Ala	Lys 385	Pro	Ala	Leu	Lys	
gag att 1315	gcg	gat	act	tcc	gtg	aac	cac	сса	ttc	ctc	gac	gag	gtt	ttg	
Glu Ile 390	Ala	Asp	Thr	Ser 395	Val	Asn	His	Pro	Phe 400	Leu	Asp	Glu	Val	Leu 405	
cac atc	atg	ggc	att	tcc	cgc	gac	gag	atc	gat	ctg	gcg	gat	cag	gaa	
His Ile	Met	Gly	Ile 410	Ser	Arg	Asp	Glu	Ile 415	Asp	Leu	Ala	Asp	Gln 420	Glu	

gac ggc act ttc cac cgc gtt cca ttg acc aat gcc taaagattcg 1409 Asp Gly Thr Phe His Arg Val Pro Leu Thr Asp Ala

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<211> 433

<212> PRT

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Thr Val Ser Gly Lys Asp Arg Pro Gly Val Thr Ala Ala Phe Phe Arg 35 40 45

Val Leu Ser Ala Asn Gln Val Gln Val Leu Asp Val Glu Gln Ser Met 50 60

Phe Arg Gly Phe Leu Asn Leu Ala Ala Phe Val Gly Ile Ala Pro Glu 65 70 75 80

Arg Val Glu Thr Val Thr Gly Leu Thr Asp Thr Leu Lys Val His 85 90 95

Gly Gln Ser Val Val Glu Leu Gln Glu Thr Val Gln Ser Ser Arg  $100 \,$   $105 \,$   $110 \,$ 

Pro Arg Ser Ser His Val Val Val Leu Gly Asp Pro Val Asp Ala 115 120 125

Leu Asp Ile Ser Arg Ile Gly Gln Thr Leu Ala Asp Tyr Asp Ala Asn 130 140

Leu Lys Val Thr Val Pro Asp Val Ser Pro Gly Gly Gly Glu Ala Met 165 170 175

Arg Lys Ala Leu Ala Ala Leu Thr Ser Glu Leu Asn Val Asp Ile Ala 180 185 190

Ile Glu Arg Ser Gly Leu Leu Arg Arg Ser Lys Arg Leu Val Cys Phe
195 200 205

Asp Cys Asp Ser Thr Leu Ile Thr Gly Glu Val Ile Glu Met Leu Ala 210 215 220

Ala His Ala Gly Lys Glu Ala Glu Val Ala Ala Val Thr Glu Arg Ala 225 230 235 240

Met Arg Gly Glu Leu Asp Phe Glu Glu Ser Leu Arg Glu Arg Val Lys

Ala Leu Ala Gly Leu Asp Ala Ser Val Ile Asp Glu Val Ala Ala Ala Ile Glu Leu Thr Pro Gly Ala Arg Thr Thr Ile Arg Thr Leu Asn Arg 280 Met Gly Tyr Gln Thr Ala Val Val Ser Gly Gly Phe Ile Gln Val Leu Glu Gly Leu Ala Glu Glu Leu Glu Leu Asp Tyr Val Arg Ala Asn Thr 310 Leu Glu Ile Val Asp Gly Lys Leu Thr Gly Asn Val Thr Gly Lys Ile 330 Val Asp Arg Ala Ala Lys Ala Glu Phe Leu Arg Glu Phe Ala Ala Asp Ser Gly Leu Lys Met Tyr Gln Thr Val Ala Val Gly Asp Gly Ala Asn 360 Asp Ile Asp Met Leu Ser Ala Ala Gly Leu Gly Val Ala Phe Asn Ala 375 Lys Pro Ala Leu Lys Glu Ile Ala Asp Thr Ser Val Asn His Pro Phe 385 390 395 Leu Asp Glu Val Leu His Ile Met Gly Ile Ser Arg Asp Glu Ile Asp 410 Leu Ala Asp Gln Glu Asp Gly Thr Phe His Arg Val Pro Leu Thr Asn 425 Ala <210> 155 <211> 490 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(490) <223> FRXA02479 <400> 155 atacatetea eccaatteee cataactaga caattgeeea geaacgaetg ataagtetee 60 aatgtegtgt teegegetea gacatgagae aattgttgee gtg act gaa ete ate 115 Val Thr Glu Leu Ile 1 cag aat gaa too caa gaa ato got gag otg gaa goo ggo cag cag gtt Gln Asn Glu Ser Gln Glu Ile Ala Glu Leu Glu Ala Gly Gln Gln Val 10 15

Ala Leu Arg	gaa ggt Glu Gly 25			~ -	-				-			211
gac cgc cca Asp Arg Pro 40												259
cag gtt cag Gln Val Gln 55			Glu									307
aac ttg gcg Asn Leu Ala 70												355
acc aca ggc Thr Thr Gly												403
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235

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acco	cttgg	gtg (	cgcg	cacca	ac ga	atcc	gtace	g gt:	tgaad	ccgc		ggt Gly		_	_	115
												ggt Gly				163
												gaa Glu				211
												gac Asp 50				259
-	_				_			-				ggc Gly				307
												atc Ile				355
												cct Pro				403
												gac Asp				451
												gcg Ala 130				499
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ttt	ctcga	acg (	ccc													558
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236

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tee tee etc atc gtt tte gee caa gga etc tte egg aag aaa tte tte

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Ser	Ser	Leu 120	Ile	Val	Phe	Ala	Gln 125	Gly	Leu	Phe	Arg	Lys 130	Lys	Phe	Phe	
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Tyr Gly Met Glu Leu Asn Glu Phe Asn Ala Gly Val Asp Ala Val Ala 65 70 75 80

Gly Ala Ile Glu Ser Ala Gly Ala Ile His Val Ser Ile Pro Asp Pro 85 90 95

Asp Val Pro Gln Asp Val Gly Ala Ala Ala Phe Phe Asp Val Asp Asn 100 105 110

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Arg Lys Lys Phe Phe Thr Ile Lys Glu Ile Leu Pro Val Val Trp Lys 130 135 140

Gln Val Lys Phe Lys Leu Thr Gly Ser Glu Asn Ala Asp Asp Val Ser 145 150 155 160

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Glu Leu Val Asp Leu Cys Glu Glu Ile Val Asp Gln Arg Met Ala Asp 180 185 190

Lys Met Trp Pro Gly Thr Lys Gln Leu Ala Asp Met His Ile Ala Ala 195 200 205

Gly His Gln Val Trp Leu Val Ser Ala Thr Pro Val Gln Leu Ala Gln 210 220

Ile Leu Ala Gln Arg Leu Gly Phe Thr Gly Ala Ile Gly Thr Val Ala225230235240

Glu Ala Lys Asp Gly Val Phe Thr Gly Arg Leu Val Gly Asp Ile Leu 245 250 255

His Gly Pro Gly Lys Arg His Ala Val Ala Ala Leu Ala Ser Ile Glu

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Leu	Pro 290	Met	Leu	Ser	Met	Val 295	Gly	Thr	Ala	Val	Ala 300	Val	Asn	Pro	Asp	
Ser 305	Lys	Leu	Arg	Lys	Glu 310	Ala	Glu	Thr	Arg	Gly 315	Trp	Asp	Val	Arg	Asp 320	
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ago gaa gao ttt ogg ato goo tog cat gaa ooa ato aaa gag ogg tgo
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Ser Glu Asp Phe Arg Ile Ala Ser His Glu Pro Ile Lys Glu Arg Cys
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acg tat aga aag cta acc ttt tta agt gcg cgg ttt tagggtgaga
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Thr His Glu Leu Phe Pro Gly Val Asn Pro Glu Pro Asn Arg Phe Ser
Val His Tyr Asp Thr Tyr Thr Ala Asp Lys Ser Pro Ile Ile Asp Ala
Val Asp Asn Val Ile Val Leu Thr Gly Gly Ser Gly His Ala Phe Lys
Leu Ser Pro Ala Tyr Gly Glu Leu Ala Ala Gln Arg Ala Val Gly Asn
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Gly Ile Ile Glu Glu Ser Val Thr Phe Val Asn Ala Pro Arg Ile Ala 50 55 60

Glu Glu Arg Gly Leu Asp Ile Ser Val Lys Thr Asn Ser Glu Ser Val 65 70 75 80

Thr His Arg Ser Val Leu Gln Val Lys Val Ile Thr Gly Ser Gly Ala 85 90 95

Ser Ala Thr Val Val Gly Ala Leu Thr Gly Leu Glu Arg Val Glu Lys
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Ile Thr Arg Ile Asn Gly Arg Gly Leu Asp Leu Arg Ala Glu Gly Leu 115 120 125

Asn Leu Phe Leu Gln Tyr Thr Asp Ala Pro Gly Ala Leu Gly Thr Val 130 135 140

Gly Thr Lys Leu Gly Ala Ala Gly Ile Asn Ile Glu Ala Ala Ala Leu 145 150 155 160

Thr Gln Ala Glu Lys Gly Asp Gly Ala Val Leu Ile Leu Arg Val Glu 165 170 175

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107	75														acc	
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Val Met Arg Leu Met Thr Glu Tyr Gly Asp Glu Leu Ala His Arg Ile 35 40 45

Gly Gly Pro Leu Glu Val Arg Gly Ile Ala Val Ser Asp Ile Ser Lys 50 55 60

Pro Arg Glu Gly Val Ala Pro Glu Leu Leu Thr Glu Asp Ala Phe Ala Leu Ile Glu Arg Glu Asp Val Asp Ile Val Val Glu Val Ile Gly Gly Ile Glu Tyr Pro Arg Glu Val Val Leu Ala Ala Leu Lys Ala Gly Lys 105 Ser Val Val Thr Ala Asn Lys Ala Leu Val Ala Ala His Ser Ala Glu 120 Leu Ala Asp Ala Ala Glu Ala Ala Asn Val Asp Leu Tyr Phe Glu Ala 135 Ala Val Ala Cys Ala Ile Pro Val Val Gly Pro Leu Arg Arg Ser Leu Ala Gly Asp Gln Ile Gln Ser Val Met Gly Ile Val Asn Gly Thr Thr 170 Asn Phe Ile Leu Asp Ala Met Asp Ser Thr Gly Ala Asp Tyr Ala Asp Ser Leu Ala Glu Ala Thr Arg Leu Gly Tyr Ala Glu Ala Asp Pro Thr Ala Asn Val Glu Gly His Asp Ala Ala Ser Lys Ala Ala Ile Leu Ala Cys Ile Ala Phe His Thr Arg Val Thr Ala Asp Asp Val Tyr Cys Glu Gly Ile Arg Asn Ile Asn Ala Ala Asp Ile Glu Ala Ala Gln Gln Ala Gly His Thr Ile Lys Leu Leu Ala Ile Cys Glu Lys Phe Thr Asn Lys 265 Glu Gly Lys Ser Ala Ile Ser Ala Arg Val His Pro Thr Leu Leu Pro 280 Val Ser His Pro Leu Ala Ser Val Asn Lys Ser Phe Asn Ala Ile Phe Val Glu Ala Glu Ala Ala Gly Arg Leu Met Phe Tyr Gly Asn Gly Ala 315 Gly Gly Ala Pro Thr Ala Ser Ala Val Leu Gly Asp Val Val Gly Ala Ala Arg Asn Lys Val His Gly Gly Arg Ala Pro Gly Glu Ser Thr Tyr 345 Ala Asn Leu Pro Ile Ala Asp Phe Gly Glu Thr Thr Thr Arg Tyr His 360 355 Leu Asp Met Asp Val Glu Asp Arg Val Gly Val Leu Ala Glu Leu Ala 375 Ser Leu Phe Ser Glu Gln Gly Ile Ser Leu Arg Thr Ile Arg Gln Glu

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                             40
Gly Gly Pro Leu Glu Val Arg Gly Ile Ala Val Ser Asp Ile Ser Lys
Pro Arg Glu Gly Val Ala Pro Glu Leu Leu Thr Glu Asp Ala Phe Ala
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Leu Ile Glu Arg Glu Asp Val Asp Ile Val Val Glu Val Ile Gly Gly
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                                             Met Ala Ile Glu Leu
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Asn Val Gly Arg Lys Val Thr Val Thr Val Pro Gly Ser Ser Ala Asn
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 ctc gga cct ggc ttt gac act tta ggt ttg gca ctg tcg gta tac gac
 Leu Gly Pro Gly Phe Asp Thr Leu Gly Leu Ala Leu Ser Val Tyr Asp
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	gct Ala															355
	cga Arg															403
	tct Ser															451
	gat Asp															499
	gaa Glu 135															547
	gtg Val															595
	gct Ala															643
	gtt Val															691
	act Thr															739
	gtg Val 215															787
	act Thr															835
	acc Thr															883
	ctt Leu															931
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<213> Corynebacterium glutamicum

<400> 174

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Glu Val Glu Val Phe Gly Glu Gly Gln Gly Glu Val Pro Leu Asp Gly
50 60

Ser His Leu Val Val Lys Ala Ile Arg Ala Gly Leu Lys Ala Ala Asp 65 70 75 80

Ala Glu Val Pro Gly Leu Arg Val Val Cys His Asn Asn Ile Pro Gln
85 90 95

Ser Arg Gly Leu Gly Ser Ser Ala Ala Ala Ala Val Ala Gly Val Ala
100 105 110

Ala Ala Asn Gly Leu Ala Asp Phe Pro Leu Thr Gln Glu Gln Ile Val 115 120 125

Gln Leu Ser Ser Ala Phe Glu Gly His Pro Asp Asn Ala Ala Ala Ser 130 135 140

Val Leu Gly Gly Ala Val Val Ser Trp Thr Asn Leu Ser Ile Asp Gly 145 150 155 160

Lys Ser Gln Pro Gln Tyr Ala Ala Val Pro Leu Glu Val Gln Asp Asn 165 170 175

Ile Arg Ala Thr Ala Leu Val Pro Asn Phe His Ala Ser Thr Glu Ala 180 185 190

Val Arg Arg Val Leu Pro Thr Glu Val Thr His Ile Asp Ala Arg Phe 195 200 205

Asn Val Ser Arg Val Ala Val Met Ile Val Ala Leu Gln Gln Arg Pro 210 215 220

Asp Leu Leu Trp Glu Gly Thr Arg Asp Arg Leu His Gln Pro Tyr Arg 225 230 235 240

Ala Glu Val Leu Pro Ile Thr Ser Glu Trp Val Asn Arg Leu Arg Asn 250 Arg Gly Tyr Ala Ala Tyr Leu Ser Gly Ala Gly Pro Thr Ala Met Val 265 Leu Ser Thr Glu Pro Ile Pro Asp Lys Val Leu Glu Asp Ala Arg Glu 280 Ser Gly Ile Lys Val Leu Glu Leu Glu Val Ala Gly Pro Val Lys Val 295 Glu Val Asn Gln Pro 305 <210> 175 <211> 1566 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101) . . (1543) <223> RXA00330 <400> 175 qcaacacttt agggtatcgc gtgggcgaag tcaccttttt caacatattt gagacggtgt 60 qqqqqaqtat tqtgtcaccc cttgggatag ggttatatcc gtg gac tac att tcg 115 Val Asp Tyr Ile Ser 163 acg cgt gat gcc agc cgt acc cct gcc cgc ttc agt gat att ttg ctg Thr Arg Asp Ala Ser Arg Thr Pro Ala Arg Phe Ser Asp Ile Leu Leu 10 ggc ggt cta gca cca gac ggc gga ctg tac ctg cct gca acc tac cct 211 Gly Gly Leu Ala Pro Asp Gly Gly Leu Tyr Leu Pro Ala Thr Tyr Pro 259 caa cta gat gat gcc cag ctg agt aaa tgg cgt gag gta tta gcc aac Gln Leu Asp Asp Ala Gln Leu Ser Lys Trp Arg Glu Val Leu Ala Asn 40 45 307 gaa gga tac gca gct ttg gct gct gaa gtt atc tcc ctg ttt gtt gat Glu Gly Tyr Ala Ala Leu Ala Ala Glu Val Ile Ser Leu Phe Val Asp 60 gac atc cca gta gaa gac atc aag gcg atc acc gca cgc gcc tac acc 355 Asp Ile Pro Val Glu Asp Ile Lys Ala Ile Thr Ala Arg Ala Tyr Thr tac ccg aag ttc aac agc gaa gac atc gtt cct gtc acc gaa ctc gag 403 Tyr Pro Lys Phe Asn Ser Glu Asp Ile Val Pro Val Thr Glu Leu Glu 95 gac aac att tac ctg ggc cac ctt tcc gaa ggc cca acc gct gca ttc Asp Asn Ile Tyr Leu Gly His Leu Ser Glu Gly Pro Thr Ala Ala Phe 105 110

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tto 112		ı cgt	ttc	ato	ttc	gac	ctg	ctc	ggc	cgc	gac	gcc	acc	c gc	gtc	
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355 360 365 Tyr Gly Phe Ala Ser Gly Arg Ser Thr His Ala Asp Arg Val Ala Thr 375 Ile Ala Asp Val His Ser Arg Leu Asp Val Leu Ile Asp Pro His Thr 390 395 Ala Asp Gly Val His Val Ala Arg Gln Trp Arg Asp Glu Val Asn Thr 410 Pro Ile Ile Val Leu Glu Thr Ala Leu Pro Val Lys Phe Ala Asp Thr 420 Ile Val Glu Ala Ile Gly Glu Ala Pro Gln Thr Pro Glu Arg Phe Ala Ala Ile Met Asp Ala Pro Phe Lys Val Ser Asp Leu Pro Asn Asp Thr Asp Ala Val Lys Gln Tyr Ile Val Asp Ala Ile Ala Asn Thr Ser Val 465 Lys <210> 177 <211> 1254 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1231) <223> RXN00403 tttttcagac tcgtgagaat gcaaactaga ctagacagag ctgtccatat acactggacg 60 aagttttagt cttgtccacc cagaacaggc ggttattttc atg ccc acc ctc gcg 115 Met Pro Thr Leu Ala cct tca ggt caa ctt gaa atc caa gcg atc ggt gat gtc tcc acc gaa Pro Ser Gly Gln Leu Glu Ile Gln Ala Ile Gly Asp Val Ser Thr Glu gcc gga gca atc att aca aac gct gaa atc gcc tat cac cgc tgg ggt Ala Gly Ala Ile Ile Thr Asn Ala Glu Ile Ala Tyr His Arg Trp Gly gaa tac cgc gta gat aaa gaa gga cgc agc aat gtc gtt ctc atc gaa 259 Glu Tyr Arg Val Asp Lys Glu Gly Arg Ser Asn Val Val Leu Ile Glu 40 4.5 307 cac gcc ctc act gga gat tcc aac gca gcc gat tgg tgg gct gac ttg His Ala Leu Thr Gly Asp Ser Asn Ala Ala Asp Trp Trp Ala Asp Leu 60 355 ctc ggt ccc ggc aaa gcc atc aac act gat att tac tgc gtg atc tgt

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				aat Asn												451
cgt Arg	gat Asp	cag Gln 120	gta Val	aac Asn	gcc Ala	gaa Glu	aaa Lys 125	caa Gln	ttc Phe	ctc Leu	gac Asp	gca Ala 130	ctc Leu	ggc	atc Ile	499
				gca Ala												547
				gca Ala												595
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				gcc Ala												835
				cgc Arg 250												883
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102	7			tcc												
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Leu Gly Asn Leu Leu Ala Met Ala Lys Ile Val Ser Pro Val Gly His 330 335 340

gat gct ttc ctc acc gaa agc cgc caa atg gat cgc atc gtg agg aac 1171

Asp Ala Phe Leu Thr Glu Ser Arg Gln Met Asp Arg Ile Val Arg Asn 345 350 355

ttc ttc agc ctc atc tcc cca gac gaa gac aac cct tcg acc tac atc 1219

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Val Val Leu Ile Glu His Ala Leu Thr Gly Asp Ser Asn Ala Ala Asp

Trp Trp Ala Asp Leu Gly Pro Gly Lys Ala Ile Asn Thr Asp Ile
65 70 75 80

Tyr Cys Val Ile Cys Thr Asn Val Ile Gly Gly Cys Asn Gly Ser Thr 85 90 95

Gly Pro Gly Ser Met His Pro Asp Gly Asn Phe Trp Gly Asn Arg Phe
100 105 110

Pro Ala Thr Ser Ile Arg Asp Gln Val Asn Ala Glu Lys Gln Phe Leu 115 120 125

Asp Ala Leu Gly Ile Thr Thr Val Ala Ala Val Leu Gly Gly Ser Met 130 140

Gly Gly Ala Arg Thr Leu Glu Trp Ala Ala Met Tyr Pro Glu Thr Val 145 150 155 160

Gly Ala Ala Ala Val Leu Ala Val Ser Ala Arg Ala Ser Ala Trp Gln Ile Gly Ile Gln Ser Ala Gln Ile Lys Ala Ile Glu Asn Asp His His 185 Trp His Glu Gly Asn Tyr Tyr Glu Ser Gly Cys Asn Pro Ala Thr Gly 200 Leu Gly Ala Ala Arg Arg Ile Ala His Leu Thr Tyr Arg Gly Glu Leu Glu Ile Asp Glu Arg Phe Gly Thr Lys Ala Gln Lys Asn Glu Asn Pro 230 235 Leu Gly Pro Tyr Arg Lys Pro Asp Gln Arg Phe Ala Val Glu Ser Tyr 245 Leu Asp Tyr Gln Ala Asp Lys Leu Val Gln Arg Phe Asp Ala Gly Ser 265 Tyr Val Leu Leu Thr Asp Ala Leu Asn Arg His Asp Ile Gly Arg Asp 280 Arg Gly Gly Leu Asn Lys Ala Leu Glu Ser Ile Lys Val Pro Val Leu 295 Val Ala Gly Val Asp Thr Asp Ile Leu Tyr Pro Tyr His Gln Glu 310 305 His Leu Ser Arg Asn Leu Gly Asn Leu Leu Ala Met Ala Lys Ile Val 330 325 Ser Pro Val Gly His Asp Ala Phe Leu Thr Glu Ser Arg Gln Met Asp 340 Arg Ile Val Arg Asn Phe Phe Ser Leu Ile Ser Pro Asp Glu Asp Asn Pro Ser Thr Tyr Ile Glu Phe Tyr Ile 370 <210> 179 <211> 1210 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1210) <223> FRXA00403 <400> 179 tttttcagac tcgtgagaat gcaaactaga ctagacagag ctgtccatat acactggacg 60 aagttttagt cttgtccacc cagaacagge ggttattttc atg ccc acc ctc gcg Met Pro Thr Leu Ala

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-		_			aca Thr		_	_		_				_		211
					aaa Lys											259
					gat Asp											307
					gcc Ala 75											355
					ggt Gly											403
					ttc Phe											451
					gcc Ala											499
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					atg Met 155											595
					cgc Arg											643
					att Ile				cac					ggc		691
					tgc Cys											739
					acc Thr											787
					caa Gln 235											835
aag	CCC	gac	cag	cgc	ttc	gcc	gtg	gaa	tcc	tac	ttg	gac	tac	caa	gca	883

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Gly Pro Gly Ser Met His Pro Asp Gly Asn Phe Trp Gly Asn Arg Phe 105 100 Pro Ala Thr Ser Ile Arg Asp Gln Val Asn Ala Glu Lys Gln Phe Leu 120 Asp Ala Leu Gly Ile Thr Thr Val Ala Ala Val Leu Gly Gly Ser Met 135 Gly Gly Ala Arg Thr Leu Glu Trp Ala Ala Met Tyr Pro Glu Thr Val Gly Ala Ala Ala Val Leu Ala Val Ser Ala Arg Ala Ser Ala Trp Gln Ile Gly Ile Gln Ser Ala Gln Ile Lys Ala Ile Glu Asn Asp His His Trp His Glu Gly Asn Tyr Tyr Glu Ser Gly Cys Asn Pro Ala Thr Gly Leu Gly Ala Ala Arg Arg Ile Ala His Leu Thr Tyr Arg Gly Glu Leu 215 Glu Ile Asp Glu Arg Phe Gly Thr Lys Ala Gln Lys Asn Glu Asn Pro 230 Leu Gly Pro Tyr Arg Lys Pro Asp Gln Arg Phe Ala Val Glu Ser Tyr Leu Asp Tyr Gln Ala Asp Lys Leu Val Gln Arg Phe Asp Ala Gly Ser Tyr Val Leu Leu Thr Asp Ala Leu Asn Arg His Asp Ile Gly Arg Asp Arg Gly Gly Leu Asn Lys Ala Leu Glu Ser Ile Lys Val Pro Val Leu Val Ala Gly Val Asp Thr Asp Ile Leu Tyr Pro Tyr His Gln Glu ASH Leu Gly ASH Leu Leu Ala Met Ala Lys Ile Val 330 Ser Pro Val Gly His Asp Ala Phe Leu Thr Glu Ser Arg Gln Met Asp Arg Ile Val Arg Asn Phe Phe Ser Leu Ile Ser Pro Asp Glu Asp Asn 360 Pro Ser 370 <210> 181 <211> 771 <212> DNA <213> Corynebacterium glutamicum <220>

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Ser Leu Arg

215

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Pro Thr Asp Thr Leu Tyr Gly Leu Gly Cys Asp Ala Phe Asn Asn Glu 35 40 45

Ala Val Ala Asn Leu Leu Ala Thr Lys His Arg Gly Pro Asp Met Pro 50 60

Val Pro Val Leu Val Gly Ser Trp Asp Thr Ile Gln Gly Leu Val His 65 70 75 80

Ser Tyr Ser Ala Gln Ala Lys Ala Leu Val Glu Ala Phe Trp Pro Gly 85 90 95

Gly Leu Ser Ile Ile Val Pro Gln Ala Pro Ser Leu Pro Trp Asn Leu 100 105 110

Gly Asp Thr Arg Gly Thr Val Met Leu Arg Met Pro Leu His Pro Val 115 120 125

Ala Ile Glu Leu Leu Arg Gln Thr Gly Pro Met Ala Val Ser Ser Ala 130 135 140

Asn Ile Ser Gly His Thr Pro Pro Thr Thr Val Leu Glu Ala Arg Gln 145 150 155 160

Gln Leu Asn Gln Asn Val Ala Val Tyr Leu Asp Gly Gly Glu Cys Ala 165 170 175

Leu Ala Thr Pro Ser Thr Ile Val Asp Ile Ser Gly Pro Ala Pro Lys
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S. CDOO!B 140

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Ile Glu Arg Leu Gly Ser Tyr Thr Arg Thr Val Ser Gly Gly Leu Thr 35 40 45

Leu Leu Val Pro Phe Val Asp Arg Val Arg Ala Arg Ile Asp Thr Arg 50 55 60

Glu Arg Val Val Ser Phe Pro Pro Gln Ala Val Ile Thr Gln Asp Asn 65 70 75 80

Leu Thr Val Ala Ile Asp Ile Val Val Thr Phe Gln Ile Asn Glu Pro
85 90 95

Glu Arg Ala Ile Tyr Gly Val Asp Asn Tyr Ile Val Gly Val Glu Gln 100 105 110

Ile Ser Val Ala Thr Leu Arg Asp Val Val Gly Gly Met Thr Leu Glu 115 120 125

Glu Thr Leu Thr Ser Arg Asp Val Ile Asn Arg Arg Leu Arg Gly Glu 130 135 140

Leu Asp Ala Ala Thr Thr Lys Trp Gly Leu Arg Ile Ser Arg Val Glu 145 150 155 160

Leu Lys Ala Ile Asp Pro Pro Pro Ser Ile Gl<br/>n Gl<br/>n Ser Met Glu Lys 165 170 175

Gln Met Lys Ala Asp Arg Clu Lys Arg Ala Thr Ile Leu Thr Ala Glu 180 185 190

Gly Gln Arg Glu Ala Asp Ile Lys Thr Ala Glu Gly Glu Lys Gln Ala 195 200 205

Lys Ile Leu Gln Ala Glu Gly Glu Lys His Ala Ser Ile Leu Asn Ala 210 215 220

Glu Ala Glu Arg Gln Ala Met Ile Leu Arg Ala Glu Gly Glu Arg Ala 225 230 235 240

Ala Arg Tyr Leu Gln Ala Gln Gly Glu Ala Arg Ala Ile Gln Lys Val Asn Ala Ala Ile Lys Ser Ala Lys Leu Thr Pro Glu Val Leu Ala Tyr Gln Tyr Leu Glu Lys Leu Pro Lys Ile Ala Glu Gly Asn Ala Ser Lys 280 275 Met Trp Val Ile Pro Ser Gln Phe Ser Asp Ser Leu Glu Gly Phe Ala 295 Lys Gln Phe Gly Ala Lys Asp Ala Glu Gly Val Phe Arg Tyr Glu Pro 315 305 310 Asn Thr Val Asp Glu Glu Thr Arg Asp Ile Ala Asn Ala Asp Asn Val 330 Glu Asp Trp Phe Ser Thr Glu Ser Asp Pro Glu Ile Ala Ala Val 340 345 Ala Ala Asn Ala Val Ala Asn Lys Pro Val Asp Pro Glu Pro Gly 360 Glu Ile Leu Ser Lys Lys Thr Ala Arg Arg Val Glu Pro Glu Ala Val 370 375 380 Leu Glu Ala Leu Gln Asn Gly Thr Thr Thr Gln Pro Glu Val Glu Ala 390 395 Ala Pro Pro Thr Ala Asn Phe Ala Gln Glu Phe Pro Ala Pro Gln Ala 405 410 Asn Pro Glu Asp Tyr Ser Asp Gln His Arg Glu Asn Pro Tyr Gly Asn 430 420 425

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							gcc Ala		259
							cag Gln		307
							atc Ile		355
							aac Asn		403
							att Ile 115		451
							ggt Gly		499
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							gct Ala		595
							acc Thr		643
							ggc Gly 195		691
							tcg Ser		739
							gag Glu		787
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PCT/IB00/00923 WO 01/00843

tgg aag tgg cag cat ggc gat gtc tct cgc cac acc ggc ggc gac ttg 931 Trp Lys Trp Gln His Gly Asp Val Ser Arg His Thr Gly Gly Asp Leu 270 265 gca gcg gct ctt ggc cga gtg aag gct aag acc ttc gtt atg ccc atc 979 Ala Ala Leu Gly Arg Val Lys Ala Lys Thr Phe Val Met Pro Ile 280 285 age gag gae atg tte ttt eet gtt egt gae tgt gee gea gaa eaa gea 1027 Ser Glu Asp Met Phe Phe Pro Val Arg Asp Cys Ala Ala Glu Gln Ala 300 ctc atc cca ggc agc gag ctt cga gtg atc gaa gac atc gcc ggt cac Leu Ile Pro Gly Ser Glu Leu Arg Val Ile Glu Asp Ile Ala Gly His 320 ctt ggg ctt ttt aac gtc tct gag aat tac atc cca cag atc gac aaa Leu Gly Leu Phe Asn Val Ser Glu Asn Tyr Ile Pro Gln Ile Asp Lys aat ctg aaa gag ctg ttc gag agc taaacactga tgtcaaagag cct 1170 Asn Leu Lys Glu Leu Phe Glu Ser 345 <210> 186 <211> 349 <212> PRT <213> Corynebacterium glutamicum <400> 186 Met Leu Asp Asn Ser Phe Tyr Thr Ala Glu Val Gln Gly Pro Tyr Glu 5 15 Thr Ala Ser Ile Gly Arg Leu Glu Leu Glu Gly Gly Val Ile Glu Asp Cys Trp Leu Ala Tyr Ala Thr Ala Gly Thr Leu Asn Glu Asp Lys Ser Asn Ala Ile Leu Ile Pro Thr Trp Tyr Ser Gly Thr His Gln Thr

55

Trp Phe Gln Gln Tyr Ile Gly Thr Asp His Ala Leu Asp Pro Ser Lys 65 70 75

Tyr Phe Ile Ile Ser Ile Asn Gln Ile Gly Asn Gly Leu Ser Val Ser

Pro Ala Asn Thr Ala Asp Asp Ser Ile Ser Met Ser Lys Phe Pro Asn

Val Arg Ile Gly Asp Asp Val Val Ala Gln Asp Arg Leu Leu Arg Gln 120

Glu Phe Gly Ile Thr Glu Leu Phe Ala Val Val Gly Gly Ser Met Gly 135 140

Ala Gln Gln Thr Tyr Glu Trp Ile Val Arg Phe Pro Asp Gln Val His 150 Arg Ala Ala Pro Ile Ala Gly Thr Ala Lys Asn Thr Pro His Asp Phe 170 Ile Phe Thr Gln Thr Leu Asn Glu Thr Val Glu Ala Asp Pro Gly Phe 185 Asn Gly Gly Glu Tyr Ser Ser His Glu Glu Val Ala Asp Gly Leu Arg 200 Arg Gln Ser His Leu Trp Ala Ala Met Gly Phe Ser Thr Glu Phe Trp 215 Lys Gln Glu Ala Trp Arg Arg Leu Gly Leu Glu Ser Lys Glu Ser Val Leu Ala Asp Phe Leu Asp Pro Leu Phe Met Ser Met Asp Pro Asn Thr 250 Leu Leu Asn Asn Ala Trp Lys Trp Gln His Gly Asp Val Ser Arg His Thr Gly Gly Asp Leu Ala Ala Leu Gly Arg Val Lys Ala Lys Thr Phe Val Met Pro Ile Ser Glu Asp Met Phe Pro Val Arg Asp Cys Ala Ala Glu Gln Ala Leu Ile Pro Gly Ser Glu Leu Arg Val Ile Glu 315 Asp Ile Ala Gly His Leu Gly Leu Phe Asn Val Ser Glu Asn Tyr Ile 330 Pro Gln Ile Asp Lys Asn Leu Lys Glu Leu Phe Glu Ser 345 <210> 187 <211> 1254 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(1231) <223> RXN00403 <400> 187 tttttcagac tcgtgagaat gcaaactaga ctagacagag ctgtccatat acactggacg 60 aagttttagt cttgtccacc cagaacaggc ggttattttc atg ccc acc ctc gcg 115 Met Pro Thr Leu Ala cct toa ggt caa ctt gaa atc caa gcg atc ggt gat gtc tcc acc gaa Pro Ser Gly Gln Leu Glu Ile Gln Ala Ile Gly Asp Val Ser Thr Glu 15 10

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	_				-				-	_				gac Asp	_	307
					_				_			_		atc Ile	_	355
		_				-								tcc Ser 100	_	403
		_							_			-	_	tcc Ser		451
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	-	-	-	_	_					_			-	cgc Arg		547
			-	-	_			- ·		-		-		gct Ala	_	595
	_	_		_	_	_	_	_						caa Gln 180		643
gcc	caa	att				-								ggc		691
miu	GIII	116	БуS 185	Ala	116	Giu	ASII	190	птр	птэ	пр	ure	195	Gly	ASII	
					-									gcc Ala		739
														gaa Glu		787
				_		_		-						tac Tyr	-	835
						-		-			-	_		caa Gln 260		883

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PCT/IB00/00923 WO 01/00843

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Ser Pro Val Gly His Asp Ala Phe Leu Thr Glu Ser Arg Gln Met Asp 340 345

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		acc Thr														835
		gac Asp														883
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		Phe	Leu	Thr	Glu	Ser	Arg	Gln	Met	Asp	Arg	Ile	Val	Arg	Asn	
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Tyr His Arg Trp Gly Glu Tyr Arg Val Asp Lys Glu Gly Arg Ser Asn 40 Val Val Leu Ile Glu His Ala Leu Thr Gly Asp Ser Asn Ala Ala Asp Trp Trp Ala Asp Leu Leu Gly Pro Gly Lys Ala Ile Asn Thr Asp Ile 75 Tyr Cys Val Ile Cys Thr Asn Val Ile Gly Gly Cys Asn Gly Ser Thr Gly Pro Gly Ser Met His Pro Asp Gly Asn Phe Trp Gly Asn Arg Phe 105 Pro Ala Thr Ser Ile Arg Asp Gln Val Asn Ala Glu Lys Gln Phe Leu Asp Ala Leu Gly Ile Thr Thr Val Ala Ala Val Leu Gly Gly Ser Met Gly Gly Ala Arg Thr Leu Glu Trp Ala Ala Met Tyr Pro Glu Thr Val 150 Gly Ala Ala Ala Val Leu Ala Val Ser Ala Arg Ala Ser Ala Trp Gln Ile Gly Ile Gln Ser Ala Gln Ile Lys Ala Ile Glu Asn Asp His His 185 Trp His Glu Gly Asn Tyr Tyr Glu Ser Gly Cys Asn Pro Ala Thr Gly Leu Gly Ala Ala Arg Arg Ile Ala His Leu Thr Tyr Arg Gly Glu Leu Glu Ile Asp Glu Arg Phe Gly Thr Lys Ala Gln Lys Asn Glu Asn Pro Leu Gly Pro Tyr Arg Lys Pro Asp Gln Arg Phe Ala Val Glu Ser Tyr Leu Asp Tyr Gln Ala Asp Lys Leu Val Gln Arg Phe Asp Ala Gly Ser Tyr Val Leu Leu Thr Asp Ala Leu Asn Arg His Asp Ile Gly Arg Asp Arg Gly Gly Leu Asn Lys Ala Leu Glu Ser Ile Lys Val Pro Val Leu 2.95 Val Ala Gly Val Asp Thr Asp Ile Leu Tyr Pro Tyr His Gln Glu His Leu Ser Arg Asn Leu Gly Asn Leu Leu Ala Met Ala Lys Ile Val 330 Ser Pro Val Gly His Asp Ala Phe Leu Thr Glu Ser Arg Gln Met Asp 340

Arg Ile Val Arg Asn Phe Phe Ser Leu Ile Ser Pro Asp Glu Asp Asn 360 Pro Ser 370 <210> 191 <211> 687 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(664) <223> RXS03158 <400> 191 caaagctcac cgaaggcacc aacgccaagt tggttgttga caacaccttg gcatccccat 60 acctgcagca gccactaaaa ctcggcgcac acgcaagtcc ttg cac tcc acc acc Leu His Ser Thr Thr aag tac atc gaa gga cac tcc gac gtt gtt ggc ggc ctt gtg ggt acc Lys Tyr Ile Glu Gly His Ser Asp Val Val Gly Gly Leu Val Gly Thr 10 211 aac gac cag gaa atg gac gaa gaa ctg ctg ttc atg cag ggc ggc atc Asn Asp Gln Glu Met Asp Glu Glu Leu Leu Phe Met Gln Gly Gly Ile 25 gga ccg atc cca tca gtt ttc gat gca tac ctg acc gcc cgt ggc ctc 259 Gly Pro Ile Pro Ser Val Phe Asp Ala Tyr Leu Thr Ala Arg Gly Leu 40 aag acc ctt gca gtg cgc atg gat cgc cac tgc gac aac gca gaa aag 307 Lys Thr Leu Ala Val Arg Met Asp Arg His Cys Asp Asn Ala Glu Lys 55 ate geg gaa tte etg gae tee ege eea gag gte tee ace gtg ete tae 355 Ile Ala Glu Phe Leu Asp Ser Arg Pro Glu Val Ser Thr Val Leu Tyr cca ggt ctg aag aac cac cca ggc cac gaa gtc gca gcg aag cag atg 403 Pro Gly Leu Lys Asn His Pro Gly His Glu Val Ala Ala Lys Gln Met aag ege tte gge gge atg ate tee gte egt tte gea gge gge gaa gaa 451 Lys Arg Phe Gly Gly Met Ile Ser Val Arg Phe Ala Gly Gly Glu Glu 105 gca gct aag aag ttc tgt acc tcc acc aaa ctg atc tgt ctg gcc gag 499 Ala Ala Lys Lys Phe Cys Thr Ser Thr Lys Leu Ile Cys Leu Ala Glu 120 125 the etc ggt gge gtg gaa tee etc etg gag cae dea gea ace atg ace 547 Ser Leu Gly Gly Val Glu Ser Leu Leu Glu His Pro Ala Thr Met Thr 135 140 cac cag tca gct gcc ggc tct cag ctc gag gtt ccc cgc gac ctc gtg 595

643

687

His Gln Ser Ala Ala Gly Ser Gln Leu Glu Val Pro Arg Asp Leu Val 160 155 cgc atc tcc att ggt att gaa gac att gaa gac ctg ctc gca gat gtc Arg Ile Ser Ile Gly Ile Glu Asp Ile Glu Asp Leu Leu Ala Asp Val 170 175 gag cag gcc ctc aat aac ctt tagaaactat ttggcggcaa gca Glu Gln Ala Leu Asn Asn Leu <210> 192 <211> 188 <212> PRT <213> Corynebacterium glutamicum <400> 192 Leu His Ser Thr Thr Lys Tyr Ile Glu Gly His Ser Asp Val Val Gly Gly Leu Val Gly Thr Asn Asp Gln Glu Met Asp Glu Glu Leu Leu Phe 25 Met Gln Gly Gly Ile Gly Pro Ile Pro Ser Val Phe Asp Ala Tyr Leu Thr Ala Arg Gly Leu Lys Thr Leu Ala Val Arg Met Asp Arg His Cys Asp Asn Ala Glu Lys Ile Ala Glu Phe Leu Asp Ser Arg Pro Glu Val Ser Thr Val Leu Tyr Pro Gly Leu Lys Asn His Pro Gly His Glu Val Ala Ala Lys Gln Met Lys Arg Phe Gly Gly Met Ile Ser Val Arg Phe Ala Gly Gly Glu Glu Ala Ala Lys Lys Phe Cys Thr Ser Thr Lys Leu 120 Ile Cys Leu Ala Glu Ser Leu Gly Gly Val Glu Ser Leu Glu His 135 Pro Ala Thr Met Thr His Gln Ser Ala Ala Gly Ser Gln Leu Glu Val 150 155 Pro Arg Asp Leu Val Arg Ile Ser Ile Gly Ile Glu Asp Ile Glu Asp 165 170 Leu Leu Ala Asp Val Glu Gln Ala Leu Asn Asn Leu 180 185 <210> 193 <211> 617 <212> DNA

281

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SK 5000 D 2000 - 010001212 L

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Pro Ile Pro Ser Val Phe Asp Ala Tyr Leu Thr Ala Arg Gly Leu Lys 50 55

Thr Leu Ala Val Arg Met Asp Arg His Cys Asp Asn Ala Glu Lys Ile 65 70 75 80

Ala Glu Phe Leu Asp Ser Arg Pro Glu Val Ser Thr Val Leu Tyr Pro 85 90 95

Gly Leu Lys Asn His Pro Gly His Glu Val Ala Ala Lys Gln Met Lys 100 \$105\$

Arg Phe Gly Gly Met Ile Ser Val Arg Phe Ala Gly Gly Glu Glu Ala 115 120 125

Ala Lys Lys Phe Cys Thr Ser Thr Lys Leu Ile Cys Leu Ala Glu Ser 130 135 140

Leu Gly Gly Val Glu Ser Leu Leu Glu His Pro Ala Thr Met Thr His 145 150 155 160

Gin Ser Ala Ala Gly Ser Gln Leu Glu Val Pro Arg Asp Leu Val Arg 165 170 175

Ile Ser Ile Gly Ile Glu Asp Ile Glu Asp Leu Leu Ala Asp Val Glu
180 185 190

Gln Ala Leu Asn Asn Leu 195

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Met Asn Pro Pro Ile

1 5

acg ttg tcc agc act tat gtt cat gat tca gaa aaa gct tat ggg cgc 163

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_					-		tat Tyr 45				-	_			_	259
_		_	_	_	_		act Thr				-	-				307
_	_						aat Asn					_		-	_	355
	~ ~		_	_	-		gtt Val	_	-	-			-	-		403
	_	_	_			_	gat Asp					_	_		_	451
							gat Asp 125									499
_				_	_		gtc Val	_	_	_			_			547
Arg	Gly 135 cgt	Leu	Gly	Val	Leu	Thr 140 gaa	_	Val ggt	Asp	Ala	Thr 145 att	Phe gtg	Ala	Thr	Pro tcg	547 595
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ctt Leu 150 gca Ala gtg Val gat Asp	Gly 135 cgt Arg acc Thr tgc Cys cat His ttg Leu 215	caa Gln aaa Lys aag Lys ggt Gly 200 tat Tyr	cgt Arg ctt Leu tct Ser 185 tca Ser tcc Ser	cca Pro atc Ile 170 gag Glu gtg Val ttg Leu	ttg Leu 155 ggt Gly cac His ccg Pro	Thr 140 gaa Glu gga Gly cat His gga Gly gtg Val 220 ctt	Ctt Leu cac His gcg Ala ggt Gly 205	yal ggt Gly tct Ser cag Gln 190 ctt Leu ctt Leu	gct Ala gat Asp 175 ttt Phe gaa Glu gat Asp cat	gat Asp 160 ctt Leu ctt Leu gcg Ala	Thr 145 att Ile ctt Leu gcc Ala ttt Phe gca Ala 225 tcg	Phe gtg Val ctt Leu act Thr ctt Leu 210 gaa Glu gtt	ctt Leu gga Gly cac His 195 gct Ala tcc Ser	tac Tyr gtc Val 180 cgt Arg ctc Leu	tcg ser 165 gca Ala cat His cgt Arg	595 643 691 739

255 260 250 qtc cta ccc tct gga tgt gga aac atg ttg tca ttt gag ctt gat gca Val Leu Pro Ser Gly Cys Gly Asn Met Leu Ser Phe Glu Leu Asp Ala 265 270 aca cct gaa cga act gat gag att ctc gaa agc ctg tca ctt tta acc Thr Pro Glu Arg Thr Asp Glu Ile Leu Glu Ser Leu Ser Leu Leu Thr 280 285 290 cac gcg acc agt tgg gga ggt gtg gaa aca gcc att gaa cgt cgc acc His Ala Thr Ser Trp Gly Gly Val Glu Thr Ala Ile Glu Arg Arg Thr agg cgg gat gct gaa gtg gtg gca gaa gta ccg atg act ctt tgc cgc Arg Arg Asp Ala Glu Val Val Ala Glu Val Pro Met Thr Leu Cys Arg 315 gtt tcc gta gga att gaa gac gtt gaa gat cta tgg gaa gac ctc aac Val Ser Val Gly Ile Glu Asp Val Glu Asp Leu Trp Glu Asp Leu Asn gcc tca atc gac aaa gtt ctg ggt tagaactcgt agccagtaac cag 1170 Ala Ser Ile Asp Lys Val Leu Gly 345 <210> 196 <211> 349 <212> PRT <213> Corynebacterium glutamicum <400> 196 Met Asn Pro Pro Ile Thr Leu Ser Ser Thr Tyr Val His Asp Ser Glu Lys Ala Tyr Gly Arg Asp Gly Asn Asp Gly Trp Gly Ala Phe Glu Ala Ala Met Gly Thr Leu Asp Gly Gly Phe Ala Val Ser Tyr Ser Ser Gly Leu Ala Ala Ala Thr Ser Ile Ala Asp Leu Val Pro Thr Gly Gly Thr Val Val Leu Pro Lys Ala Ala Tyr Tyr Gly Val Thr Asn Ile Phe Ala Arg Met Glu Ala Arg Gly Arg Leu Lys Val Arg Thr Val Asp Ala Asp Asn Thr Glu Glu Val Ile Ala Ala Ala Gln Gly Ala Asp Val Val Trp 105

Val Glu Ser Ile Ala Asn Pro Thr Met Val Val Ala Asp Ile Pro Ala 115 120 125

Ile Val Asp Gly Val Arg Gly Leu Gly Val Leu Thr Val Val Asp Ala 135 Thr Phe Ala Thr Pro Leu Arg Gln Arg Pro Leu Glu Leu Gly Ala Asp 150 145 155 Ile Val Leu Tyr Ser Ala Thr Lys Leu Ile Gly Gly His Ser Asp Leu 170 Leu Leu Gly Val Ala Val Cys Lys Ser Glu His His Ala Gln Phe Leu 180 Ala Thr His Arg His Asp His Gly Ser Val Pro Gly Gly Leu Glu Ala Phe Leu Ala Leu Arg Gly Leu Tyr Ser Leu Ala Val Arg Leu Asp Arg Ala Glu Ser Asn Ala Ala Glu Leu Ser Arg Arg Leu Asn Ala His Pro Ser Val Thr Arg Val Asn Tyr Pro Gly Leu Pro Asp Asp Pro Gln His 250 Glu Lys Ala Val Arg Val Leu Pro Ser Gly Cys Gly Asn Met Leu Ser Phe Glu Leu Asp Ala Thr Pro Glu Arg Thr Asp Glu Ile Leu Glu Ser Leu Ser Leu Leu Thr His Ala Thr Ser Trp Gly Gly Val Glu Thr Ala Ile Glu Arg Arg Thr Arg Arg Asp Ala Glu Val Val Ala Glu Val Pro 310 Met Thr Leu Cys Arg Val Ser Val Gly Ile Glu Asp Val Glu Asp Leu 330 Trp Glu Asp Leu Asn Ala Ser Ile Asp Lys Val Leu Gly 345

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Leu Ser Phe Asp Pro
1

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tgt Cys 230	Ser	tgc Cys	agg Arg	gcg Ala	gca Ala 235	Ser	gac Asp	cga Arg	tcc Ser	cat His	Gln	ttt Phe	tcg Ser	atg Met	cat His 245	835
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Gly Tyr Glu Tyr Thr Arg Val Gly Asn Pro Thr Ile Val Ala Leu Glu
Gln Thr Val Ala Ala Leu Glu Gly Ala Lys Tyr Gly Arg Ala Phe Ser
Ser Gly Met Ala Ala Thr Asp Ile Leu Phe Arg Ile Ile Leu Lys Pro
Gly Asp His Ile Val Leu Gly Asn Asp Ala Tyr Gly Gly Thr Tyr Arg
Leu Ile Asp Thr Val Phe Thr Ala Trp Gly Val Glu Tyr Thr Val Val
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Asp Thr Ser Val Val Glu Glu Val Lys Ala Ala Ile Lys Asp Asn Thr
Lys Leu Ile Trp Val Glu Thr Pro Thr Asn Pro Ala Leu Gly Ile Thr
Asp Ile Glu Ala Val Ala Lys Leu Thr Glu Gly Thr Asn Ala Lys Leu
                                    170
                                    Tyr Leu Gin Gin Pro Leu Lys
Leu Gly Ala His Ala Ser Pro Cys Thr Pro Pro Pro Ser Thr Ser Lys
                            200
Asp Thr Pro Thr Leu Leu Ala Ala Leu Trp Val Pro Thr Thr Arg Lys
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Gln Phe Ser Met His Thr
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BESTOCHE -WO 010081342 I >

703

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cag Gln	ggt Gly	gaa Glu	acg Thr 25	ggc Gly	gat Asp	ctt Leu	ctc Leu	cat His 30	att Ile	cct Pro	cag Gln	ctt Leu	ccg Pro 35	gcg Ala	cga Arg	211
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gat	gtg	gct	cgg	ggt	gcg	ggc	gcc	gat	act	gtg	, cag	att	tcc	atg	gat	787

Aso Val Ala Arg Gly Ala Gly Ala Asp Thr Val Gln Ile Ser Met Asp 220 caa gtc cgt gga aat gaa cat ttg gat ggt ttt ggt gaa acc atc acc 835 Gln Val Arg Gly Asn Glu His Leu Asp Gly Phe Gly Glu Thr Ile Thr 240 883 agt gga att cgt ctt ggt ttg ggc att acg aca gga aaa gat gtc gta Ser Gly Ile Arg Leu Gly Leu Gly Ile Thr Thr Gly Lys Asp Val Val 250 255 931 gat gaa ctg ctc gag cga ccg cgg caa aag gcc gtt gag gta gca cgc Asp Glu Leu Leu Glu Arg Pro Arg Gln Lys Ala Val Glu Val Ala Arg 270 265 979 ttt ttt gat cgt tta ggt gtg ggc cga aac tat ctc gtg gat gct gtt Phe Phe Asp Arg Leu Gly Val Gly Arg Asn Tyr Leu Val Asp Ala Val 285 280 gat att cat ccg ggt gag gat ttg gtg cag ggg acc atc acc gag gcc 1027 Asp Ile His Pro Gly Glu Asp Leu Val Gln Gly Thr Ile Thr Glu Ala gcg cag gct tat cgc atg gcc cgg gtg atg tcg gag atg ttg tcg aag Ala Gln Ala Tyr Arg Met Ala Arg Val Met Ser Glu Met Leu Ser Lys 320 315 gat toa tgc gac ctt taaggettta ceggegetgg gtg 1113 Asp Ser Cys Asp Leu <210> 202 <211> 330 <212> PRT <213> Corynebacterium glutamicum <400> 202 Leu Gly Ala Tyr Gly Leu Gly Glu Leu Pro Gly Lys Ser Ala Ala Glu Ala Ala Asp Ile Ile Gln Gly Glu Thr Gly Asp Leu Leu His Ile Pro Gln Leu Pro Ala Arg Gly Leu Gly Ala Asp Leu Ile Gly Arg Thr Val Gly Leu Leu Asp Met Ile Asn Val Asp Arg Gly Ala Arg Ser Trp Val Met Ser Thr Arg Pro Ser Arg Leu Thr His Leu Thr Gly Asp Phe Leu 70 Asp Met Asp Leu Asp Ala Cys Glu Glu Thr Trp Gly Thr Gly Val Asp Lys Leu Lys Ile Gln Val Ala Gly Pro Trp Thr Leu Gly Ala Arg Ile 105

Glu Leu Ala Asn Gly His Arg Val Leu Ser Asp Arg Gly Ala Met Arg 120 115 Asp Leu Thr Gln Ala Leu Ile Ala Gly Ile Asp Ala His Ala Arg Lys Val Ala Gly Arg Phe Arg Ala Glu Val Gln Val Gln Ile Asp Glu Pro 150 Glu Leu Lys Ser Leu Ile Asp Gly Ser Leu Pro Gly Thr Ser Thr Phe Asp Ile Ile Pro Ala Val Asn Val Ala Asp Ala Ser Glu Arg Leu Gln 185 Gln Val Phe Ser Ser Ile Glu Gly Pro Thr Tyr Leu Asn Leu Thr Gly 200 Gln Ile Pro Thr Trp Asp Val Ala Arg Gly Ala Gly Ala Asp Thr Val 215 Gln Ile Ser Met Asp Gln Val Arg Gly Asn Glu His Leu Asp Gly Phe 230 Gly Glu Thr Ile Thr Ser Gly Ile Arg Leu Gly Leu Gly Ile Thr Thr Gly Lys Asp Val Val Asp Glu Leu Leu Glu Arg Pro Arg Gln Lys Ala 265 Val Glu Val Ala Arg Phe Phe Asp Arg Leu Gly Val Gly Arg Asn Tyr 280 Leu Val Asp Ala Val Asp Ile His Pro Gly Glu Asp Leu Val Gln Gly Thr Ile Thr Glu Ala Ala Gln Ala Tyr Arg Met Ala Arg Val Met Ser 315 310 Glu Met Leu Ser Lys Asp Ser Cys Asp Leu 325 <210> 203 <211> 623 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (1)..(600) <223> RXN00402 <400> 203 act gac gaa aag gat gga aag cca gta ttg ccc tac ttc gtc act cca 48 Thr Asp Glu Lys Asp Gly Lys Pro Val Leu Pro Tyr Phe Val Thr Pro 1 gat gct gct tac cac gga ttg aag tac gca gac ctt ggt gca cca gcc Asp Ala Ala Tyr His Gly Leu Lys Tyr Ala Asp Leu Gly Ala Pro Ala

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Leu Asn Leu Ala Gly Ala Gly Asp His Ile Val Thr Ser Pro Arg Leu

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Met Ser Thr Ser Val

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ENSURED AND PROPERTY

cag ctg gcc gct atg cct ttg ttt gag cgt ttg gca cag cgc atc atc 2131 Gln Leu Ala Ala Met Pro Leu Phe Glu Arg Leu Ala Gln Arg Ile Ile 665 670 gac ggc gat aag aat ggc ctt gag gat gat ctg gaa gca ggc atg aag 2179 Asp Gly Asp Lys Asn Gly Leu Glu Asp Asp Leu Glu Ala Gly Met Lys 680 685 690 gag aag tot oot att gog atc atc aac gag gac ott otc aac ggc atg 2227 Glu Lys Ser Pro Ile Ala Ile Ile Asn Glu Asp Leu Leu Asn Gly Met 700 aag acc gtg ggt gag ctg ttt ggt tcc gga cag atg cag ctg cca ttc 2275 Lys Thr Val Gly Glu Leu Phe Gly Ser Gly Gln Met Gln Leu Pro Phe 720 gtg ctg caa tcg gca gaa acc atg aaa act gcg gtg gcc tat ttg gaa 2323 Val Leu Gln Ser Ala Glu Thr Met Lys Thr Ala Val Ala Tyr Leu Glu 735 ccg ttc atg gaa gag gaa gca gaa gct acc gga tct gcg cag gca gag 2371 Pro Phe Met Glu Glu Glu Ala Glu Ala Thr Gly Ser Ala Gln Ala Glu 745 750 ggc aag ggc aaa atc gtc gtg gcc acc gtc aag ggt gac gtg cac gat 2419 Gly Lys Gly Lys Ile Val Val Ala Thr Val Lys Gly Asp Val His Asp 760 765 atc ggc aag aac ttg gtg gac atc att ttg tcc aac aac ggt tac gac 2467 Ile Gly Lys Asn Leu Val Asp Ile Ile Leu Ser Asn Asn Gly Tyr Asp 775 780 785 gtg gtg aac ttg ggc atc aag cag cca ctg tcc gcc atg ttg gaa gca 2515 vai vai Asn Leu Giy lle Lys Gln Pro Leu Ser Ala Met Leu Glu Ala 790 795 800 gcg gaa gaa cac aaa gca gac gtc atc ggc atg tcg gga ctt ctt gtg 2563 Ala Glu Glu His Lys Ala Asp Val Ile Gly Met Ser Gly Leu Leu Val 810 815 aag too acc gtg gtg atg aag caa acc atc agc gac 2599 Lys Ser Thr Val Val Met Lys Gln Thr Ile Ser Asp 825 830

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<400> 212

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Met Gly Thr Gln Leu Gln Gly Phe Asp Leu Asp Val Glu Lys Asp Phe 35 40 45

Leu Asp Leu Glu Gly Cys Asn Glu Ile Leu Asn Asp Thr Arg Pro Asp 50 55 60

Val Leu Arg Gln Ile His Arg Ala Tyr Phe Glu Ala Gly Ala Asp Leu 65 70 75 80

Val Glu Thr Asn Thr Phe Gly Cys Asn Leu Pro Asn Leu Ala Asp Tyr

85 90 95

Asp Ile Ala Asp Arg Cys Arg Glu Leu Ala Tyr Lys Gly Thr Ala Val 100 105 110

Ala Arg Glu Val Ala Asp Glu Met Gly Pro Gly Arg Asn Gly Met Arg 115 120 125

Arg Phe Val Val Gly Ser Leu Gly Pro Gly Thr Lys Leu Pro Ser Leu 130 135 140

Gly His Ala Pro Tyr Ala Asp Leu Arg Gly His Tyr Lys Glu Ala Ala 145 \$150\$ 155 \$160\$

Leu Gly Ile Ile Asp Gly Gly Gly Asp Ala Phe Leu Ile Glu Thr Ala 165 170 175

Gln Asp Leu Gln Val Lys Ala Ala Val His Gly Val Gln Asp Ala 180 185 190

Met Ala Glu Leu Asp Thr Phe Leu Pro Ile Ile Cys His Val Thr Val 195 200 205

Glu Thr Thr Gly Thr Met Leu Met Gly Ser Glu Ile Gly Ala Ala Leu 210 215 220

Thr Ala Leu Gln Pro Leu Gly Ile Asp Met Ile Gly Leu Asn Cys Ala 225 230 235 240

Thr Gly Pro Asp Glu Met Ser Glu His Leu Arg Tyr Leu Ser Lys His 245 250 255

Ala Asp Ile Pro Val Ser Val Met Pro Asn Ala Gly Leu Pro Val Leu 260 270

Gly Lys Asn Gly Ala Glu Tyr Pro Leu Glu Ala Glu Asp Leu Ala Gl<br/>n 275 280 285

Ala Leu Ala Gly Phe Val Ser Glu Tyr Gly Leu Ser Met Val Gly Gly 290 295 300

Cys Cys Gly Thr Thr Pro Glu His Ile Arg Ala Val Arg Asp Ala Val 305 310 315 320

Val Gly Val Pro Glu Gln Glu Thr Ser Thr Leu Thr Lys Ile Pro Ala 325 330 Gly Pro Val Glu Gln Ala Ser Arg Glu Val Glu Lys Glu Asp Ser Val 340 345 350 Ala Ser Leu Tyr Thr Ser Val Pro Leu Ser Gln Glu Thr Gly Ile Ser 360 Met Ile Gly Glu Arg Thr Asn Ser Asn Gly Ser Lys Ala Phe Arg Glu 370 375 380 Ala Met Leu Ser Gly Asp Trp Glu Lys Cys Val Asp Ile Ala Lys Gln 390 Gln Thr Arg Asp Gly Ala His Met Leu Asp Leu Cys Val Asp Tyr Val 410 Gly Arg Asp Gly Thr Ala Asp Met Ala Thr Leu Ala Ala Leu Leu Ala Thr Ser Ser Thr Leu Pro Ile Met Ile Asp Ser Thr Glu Pro Glu Val 440 Ile Arg Thr Gly Leu Glu His Leu Gly Gly Arg Ser Ile Val Asn Ser Val Asn Phe Glu Asp Gly Asp Gly Pro Glu Ser Arg Tyr Gln Arg Ile Met Lys Leu Val Lys Gln His Gly Ala Ala Val Val Ala Leu Thr Ile 490 Asp Glu Glu Gly Gln Ala Arg Thr Ala Glu His Lys Val Arg Ile Ala 500 505 510 Lys Arg Leu Ile Asp Asp Ile Thr Gly Ser Tyr Gly Leu Asp Ile Lys 520 Asp Ile Val Val Asp Cys Leu Thr Phe Pro Ile Ser Thr Gly Gln Glu 530 535 540 550 555 Lys Lys Leu Tyr Pro Glu Ile His Thr Thr Leu Gly Leu Ser Asn Ile 565 570 Ser Phe Gly Leu Asn Pro Ala Ala Arg Gln Val Leu Asn Ser Val Phe Leu Asn Glu Cys Ile Glu Ala Gly Leu Asp Ser Ala Ile Ala His Ser 600 Ser Lys Ile Leu Pro Met Asn Arg Ile Asp Asp Arg Gln Arg Glu Val 615 Ala Leu Asp Met Val Tyr Asp Arg Arg Thr Glu Asp Tyr Asp Pro Leu 635 630 Gln Glu Phe Met Gln Leu Phe Glu Gly Val Ser Ala Ala Asp Ala Lys

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Glu	Ala 690	Gly	Met	Lys	Glu	Lys 695	Ser	Pro	Ile	Ala	Ile 700	Ile	Asn	Glu	Asp	
Leu 705	Leu	Asn	Gly	Met	Lys 710	Thr	Val	Gly	Glu	Leu 715	Phe	Gly	Ser	Gly	Gln 720	
Met	Gln	Leu	Pro	Phe 725	Val	Leu	Gln	Ser	Ala 730	Glu	Thr	Met	Lys	Thr 735	Ala	
Val	Ala	Tyr	Leu 740	Glu	Pro	Phe	Met	Glu 745	Glu	Glu	Ala	Glu	Ala 750	Thr	Gly	
Ser	Ala	Gln 755	Ala	Glu	Gly	Lys	Gly 760	Lys	Ile	Val	Val	Ala 765	Thr	Val	Lys	
Gly	Asp 770	Val	His	Asp	Ile	Gly 775	Lys	Asn	Leu	Val	Asp 780	Ile	Ile	Leu	Ser	
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Ala	Met	Leu	Glu	Ala 805	Ala	Glu	Glu	His	Lys 810	Ala	Asp	Val	Ile	Gly 815	Met	
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ttg	gca	aac	cat	gtg	r ttg	ato	: ggc	gac	ggc	gcc	atg	ggc	acc	cag	ctc	213

Leu	Ala	Asn	His 25	Val	Leu	Ile	Gly	Asp 30	Gly	Ala	Met	Gly	Thr 35	Gln	Leu	
	ggc Gly		_	_	-		-	_	_			_	_			259
	aat Asn 55															307
	cgc Arg	-				-		_	_	_	_	-				355
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-	gag Glu			_	~ ~	_			-		_		-	•		499
	ctg Leu 135				_	_			_	_			-	_		547
	gat Asp															595
	ggt Gly															643
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	ttc Phe	_				_		_		-						739
-	ctc Leu 215	_					-						-	-		787
_	ggt Gly		_				-		_	_				_		835
	agc Ser															883
_	gtg Val	_			_				-	-				-	_	931

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00000

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315

310

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DESCRIPTION OF CHARMAN !

645 650 Asp Ala Arg Ala Glu Gln Leu Ala Ala Met Pro Leu Phe Glu Arg Leu 665 Ala Gln Arg Ile Ile Asp Gly Asp Lys Asn Gly Leu Glu Asp Asp Leu Glu Ala Gly Met Lys Glu Lys Ser Pro Ile Ala Ile Ile Asn Glu Asp 695 Leu Leu Asn Gly Met Lys Thr Val Gly Glu Leu Phe Gly Ser Gly Gln Met Gln Leu Pro Phe Val Leu Gln Ser Ala Glu Thr Met Lys Thr Ala 725 730 Val Ala Tyr Leu Glu Pro Phe Met Glu Glu Ala Glu Ala Thr Gly Ser Ala Gln Ala Glu Gly Lys Gly Lys Ile Val Val Ala Thr Val Lys 760 Gly Asp Val His Asp Ile Gly Lys Asn Leu Val Asp Ile Ile Leu Ser Asn Asn Gly Tyr Asp Val Val Asn Leu Gly Ile Lys Gln Pro Leu Ser 790 795 Ala Met Leu Glu Ala Ala Glu Glu His Lys Ala Asp Val Ile Gly Met 805 810 Ser Gly Leu Leu Val Lys Ser Thr Val Val 820 <210> 215 <211> 621 <212> DNA <213> Corynebacterium glutamicum <220> <222> (101)..(598) <223> RXN03074 <400> 215 tttgtgggca atctggtttt ttcgtaattg tgtgggatga atctcttaaa aattcacatt 60 tagcaggaca agcatactgt tttagttcta tgctgtgggc atg act caa agt gct Met Thr Gln Ser Ala cca gaa ttc att gcc acc gca gac ctc gta gac atc atc ggc gac aac Pro Glu Phe Ile Ala Thr Ala Asp Leu Val Asp Ile Ile Gly Asp Asn gcg caa tca tgc gac act cag ttt caa aac ctt gga ggt gcc aca gaa Ala Gln Ser Cys Asp Thr Gln Phe Gln Asn Leu Gly Gly Ala Thr Glu 25 30

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_	aaa Lys 55			_	_											307
	ggc Gly															355
	ctt Leu			_						-		-			_	403
	cga Arg															451
	gga Gly			_										_		499
	gta Val 135															547
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Ile	Ile	Gly	Asp 20	Asn	Ala	Gln	Ser	Cys 25	Asp	Thr	Gln	Phe	Gln 30	Asn	Leu	
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Gly Val Leu Val Ile Asp Gly Asp Ala Ser Val His Thr Ala Leu Val

Gly Asp Ile Ile Ala Gly Leu Gly Lys Asp His Gly Trp Ser Gly Val

Ile Val Asn Gly Ala Ile Arg Asp Ser Ala Val Ile Gly Thr Met Thr 105 100 Phe Gly Cys Lys Ala Leu Gly Thr Asn Pro Arg Lys Ser Thr Lys Thr 120 115 Gly Ser Gly Glu Arg Asp Val Val Val Ser Ile Gly Gly Ile Asp Phe 135 Ile Pro Gly His Tyr Val Tyr Ala Asp Ser Asp Gly Ile Ile Val Thr 150 Glu Ala Pro Ile Lys Gln <210> 217 <211> 621 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(598) <223> FRXA02906 <400> 217 tttgtgggca atctggtttt ttcgtaattg tgtgggatga atctcttaaa aattcacatt 60 tagcaggaca agcatactgt tttagttcta tgctgtgggc atg act caa agt gct Met Thr Gln Ser Ala cca gaa ttc att gcc acc gca gac ctc gta gac atc atc ggc gac aac Pro Glu Phe Ile Ala Thr Ala Asp Leu Val Asp Ile Ile Gly Asp Asn gcg caa tca tgc gac act cag ttt caa aac ctt gga ggt gcc aca gaa Ala Gln Ser Cys Asp Thr Gln Phe Gln Asn Leu Gly Gly Ala Thr Glu 30 ttc cac gga ata ata acc acc gtg aaa tgc ttc caa gac aac gcc ctc 259 45 307 ctg aaa tcc atc ctg agc gag gat aat cct ggg gga gtg ctg gtt atc Leu Lys Ser Ile Leu Ser Glu Asp Asn Pro Gly Gly Val Leu Val Ile 60 gat ggc gac gca tcc gtg cac acc gcg cta gtt ggc gac atc att gca Asp Gly Asp Ala Ser Val His Thr Ala Leu Val Gly Asp Ile Ile Ala 75 403 gga ctt gga aaa gat cat ggt tgg tcc gga gta att gtc aac gga gca Gly Leu Gly Lys Asp His Gly Trp Ser Gly Val Ile Val Asn Gly Ala 95 att cga gac tcc gca gtc atc ggc acc atg acc ttt ggt tgt aaa gcc Ile Arg Asp Ser Ala Val Ile Gly Thr Met Thr Phe Gly Cys Lys Ala 105 110

0100843A2 I

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<213> Corynebacterium glutamicum

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Gly Gly Ala Thr Glu Phe His Gly Ile Ile Thr Thr Val Lys Cys Phe \$35\$

Gln Asp Asn Ala Leu Leu Lys Ser Ile Leu Ser Glu Asp Asn Pro Gly 50 60

Gly Val Leu Val Ile Asp Gly Asp Ala Ser Val His Thr Ala Leu Val 65 70 75 80

Gly Asp Ile Ile Ala Gly Leu Gly Lys Asp His Gly Trp Ser Gly Val 85 90 95

Ile Val Asn Gly Ala Ile Arg Asp Ser Ala Val Ile Gly Thr Met Thr  $100 \hspace{1.5cm} 105 \hspace{1.5cm} 110 \hspace{1.5cm}$ 

Phe Gly Cys Lys Ala Leu Gly Thr Asn Pro Arg Lys Ser Thr Lys Thr 115 120 125

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												aac Asn				787
												cac His				835
_				_	-		_	_		_		ggc Gly				883
												gct Ala				931
_	~ ~		~ ~	-	_	-	-	-		-	-	gac Asp 290				979
_		cag	gct	ctg	atg	gat	ggc	tac	tct	gtg	gtc	acc	gtt	gat	gag	
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_	_	aac	ctt	ggc	aac	gcc	acc	gga	cac	сса	tca	ttt	gtc	atg	tcc	
131 Leu 390		Asn	Leu	Gly	Asn 395	Ala	Thr	Gly	His	Pro 400	Ser	Phe	Val	Met	Ser 405	
aac 136		ttc	gcc	gat	cag	acc	att	gcg	cag	atc	gaa	ctg	ttc	caa	aac	

Asn Ser Phe Ala Asp Gln Thr Ile Ala Gln Ile Glu Leu Phe Gln Asn gaa gga cag tac gag aac gag gtc tac cgt ctg cct aag gtt ctc gac Glu Gly Gln Tyr Glu Asn Glu Val Tyr Arg Leu Pro Lys Val Leu Asp gaa aag gtg gca cgc atc cac gtt gag gct ctc ggc ggt cag ctc acc 1459 Glu Lys Val Ala Arg Ile His Val Glu Ala Leu Gly Gly Gln Leu Thr gaa ctg acc aag gag cag gct gag tac atc ggc gtt gac gtt gca ggc 1507 Glu Leu Thr Lys Glu Gln Ala Glu Tyr Ile Gly Val Asp Val Ala Gly cca ttc aag ccg gag cac tac cgc tac taatgattgt cagcattgag Pro Phe Lys Pro Glu His Tyr Arg Tyr gga 1557 <210> 220 <211> 478 <212> PRT <213> Corynebacterium glutamicum <400> 220 Met Ala Gln Val Met Asp Phe Lys Val Ala Asp Leu Ser Leu Ala Glu Ala Gly Arg His Gln Ile Arg Leu Ala Glu Tyr Glu Met Pro Gly Leu 25 Met Gln Leu Arg Lys Glu Phe Ala Asp Glu Gln Pro Leu Lys Gly Ala Arg Ile Ala Gly Ser Ile His Met Thr Val Gln Thr Ala Val Leu Glu Thr Leu Thr Ala Leu Gly Ala Glu Val Arg Trp Ala Ser Cys Asn Ile Phe Ser Thr Gln Asp Glu Ala Ala Ala Ile Val Val Gly Ser Gly Thr Val Glu Glu Pro Ala Gly Val Pro Val Phe Ala Trp Lys Gly 100 Glu Ser Leu Glu Glu Tyr Trp Trp Cys Ile Asn Gln Ile Phe Ser Trp 120 Gly Asp Glu Leu Pro Asn Met Ile Leu Asp Asp Gly Gly Asp Ala Thr 130 135 Met Ala Val Ile Arg Gly Arg Glu Tyr Glu Gln Ala Gly Leu Val Pro

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Leu	Arg	Glu	Val 180	Leu	Ala	Ala	Glu	Pro 185	Gly	Lys	Trp	Gly	Lys 190	Ile	Ala
Glu	Ala	Val 195	Lys	Gly	Val	Thr	Glu 200	Glu	Thr	Thr	Thr	Gly 205	Val	His	Arg
Leu	Tyr 210	His	Phe	Ala	Glu	Glu 215	Gly	Val	Leu	Pro	Phe 220	Pro	Ala	Met	Asn
Val 225	Asn	Asp	Ala	Val	Thr 230	Lys	Ser	Lys	Phe	Asp 235	Asn	Lys	Tyr	Gly	Thr 240
Arg	His	Ser	Leu	Ile 245	Asp	Gly	Ile	Asn	Arg 250	Ala	Thr	Asp	Met	Leu 255	Met
Gly	Gly	Lys	Asn 260	Val	Leu	Val	Cys	Gly 265	Tyr	Gly	Asp	Val	Gly 270	Lys	Gly
Суѕ	Ala	Glu 275	Ala	Phe	Asp	Gly	Gln 280	Gly	Ala	Arg	Val	Lys 285	Val	Thr	Glu
Ala	Asp 290	Pro	Ile	Asn	Ala	Leu 295	Gln	Ala	Leu	Met	Asp 300	Gly	Tyr	Ser	Val
Val 305	Thr	Val	Asp	Glu	Ala 310	Ile	Glu	Asp	Ala	Asp 315	Ile	Val	Ile	Thr	Ala 320
Thr	Gly	Asn	Lys	Asp 325	Ile	Ile	Ser	Phe	Glu 330	Gln	Met	Leu	Lys	Met 335	Lys
Asp	His	Ala	Leu 340	Leu	Gly	Asn	Ile	Gly 345	His	Phe	Asp	Asn	Glu 350	Ile	Asp
Met	His	Ser 355	Leu	Leu	His	Arg	Asp 360	Asp	Val	Thr	Arg	Thr 365	Thr	Ile	Lys
Pro	Gln 370	Val	Asp	Glu	Phe	Thr 375	Phe	Ser	Thr	Gly	Arg 380	Ser	Ile	Ile	Val
Leu 385	Ser	Glu	Gly	Arg	Leu 390	Leu	Asn	Leu	Gly	Asn 395	Ala	Thr	Gly	His	Pro 400
Ser	Phe	Val	Met	Ser 405		Ser	Phe	Ala	Asp 410	Gln	Thr	Ile	Ala	Gln 415	Ile
Glu	Leu	Phe	Gln 420		Glu	Gly	Gln	Tyr 425	Glu	Asn	Glu	Val	Tyr 430	Arg	Leu
Pro	Lys	Val 435	Leu	Asp	Glu	Lys	Val 440	Ala	Arg	Ile	His	Val 445	Glu	Ala	Leu
Gly	Gly 450		Leu	Thr	Glu	Leu 455	Thr	Lys	Glu	Gln	Ala 460		Tyr	Ile	Gly
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gct gag tac atc ggc gtt gac gtt gca ggc cca ttc aag ccg gag cac
                                                                   96
Ala Glu Tyr Ile Gly Val Asp Val Ala Gly Pro Phe Lys Pro Glu His
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                                                                   128
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Tyr Arg Tyr
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Tyr Arg Tyr
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                                            Met Ala Gln Val Met
gac ttc aag gtt gcc gat ctt tca cta gca gag gca gga cgt cac cag
Asp Phe Lys Val Ala Asp Leu Ser Leu Ala Glu Ala Gly Arg His Gln
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gaa Glu	ttc Phe	gca Ala 40	gac Asp	gag Glu	cag Gln	cct Pro	ttg Leu 45	aag Lys	ggc Gly	gcc Ala	cga Arg	att Ile 50	gct Ala	ggt Gly	tct Ser	259
														act Thr		307
ttg Leu 70	ggc Gly	gct Ala	gag Glu	gtt Val	cgt Arg 75	tgg Trp	gct Ala	tcc Ser	tgc Cys	aac Asn 80	att Ile	ttc Phe	tcc Ser	acc Thr	cag Gln 85	355
gat Asp	gag Glu	gct Ala	gca Ala	gcg Ala 90	gct Ala	atc Ile	gtt Val	gtc Val	ggc Gly 95	tcc Ser	ggc Gly	acc Thr	gtc Val	gaa Glu 100	gag Glu	403
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tac Tyr	tgg Trp	tgg Trp 120	tgc Cys	atc Ile	aac Asn	cag Gln	atc Ile 125	ttc Phe	agc Ser	tgg Trp	ggc Gly	gat Asp 130	gag Glu	ctg Leu	cca Pro	499
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ggt Gly 150	cgc Arg	gaa Glu	tac Tyr	gag Glu	cag Gln 155	gct Ala	ggt Gly	ctg Leu	gtt Val	cca Pro 160	cca Pro	gca Ala	gag Glu	gcc Ala	aac Asn 165	595
gat Asp	tcc Ser	gat Asp	gag Glu	tac Tyr 170	atc Ile	gca Ala	ttc Phe	ttg Leu	ggc Gly 175	atg Met	ctg Leu	cgt Arg	gag Glu	gtt Val 180	ctt Leu	643
gct Ala	gca Ala	gag Glu	cct Pro 185	ggc Gly	aag Lys	tgg Trp	ggc Gly	aag Lys 190	atc Ile	gct Ala	gag Glu	gcc Ala	gtt Val 195	aag Lys	ggt Gly	691
gtc Val	acc Thr	gag Glu 200	gaa Glu	acc Thr	acc Thr	acc Thr	ggt Gly 205	gtg Val	cac His	cgc Arg	ctg Leu	tac Tyr 210	His	ttc Phe	gct Ala	739
gaa Glu	gaa Glu 215	ggc	gtg Val	ctg Leu	cct Pro	ttc Phe 220	Pro	gcg Ala	atg Met	aac Asn	gtc Val 225	aac Asn	gac Asp	gct Ala	gtc Val	787
acc Thr 230	aag Lys	tcc Ser	aag Lys	ttt Phe	gat Asp 235	Asn	aag Lys	tac Tyr	ggc Gly	acc Thr 240	Arg	cac His	tcc Ser	ctg Leu	atc Ile 245	835
gac Asp	ggc Gly	atc Ile	aac Asn	cgc Arg 250	Ala	act Thr	gac Asp	atg Met	ctc Leu 255	Met	ggc	ggc Gly	aag Lys	aac Asn 260	Val	883

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Asp Gly Gln Gly Ala Arg Val Lys Val Thr Glu Ala Asp Pro Ile Asn
gct ctt cag gct ctg atg gat ggc tac tct gtg gtc acc gtt gat gag
1027
Ala Leu Gln Ala Leu Met Asp Gly Tyr Ser Val Val Thr Val Asp Glu
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qcc atc gag gac gcc gac atc gtg atc acc gcg acc ggc aac aag gac
Ala Ile Glu Asp Ala Asp Ile Val Ile Thr Ala Thr Gly Asn Lys Asp
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atc att tcc ttc gag cag atg ctc aag atg aag gat cac gct ctg ctg
Ile Ile Ser Phe Glu Gln Met Leu Lys Met Lys Asp His Ala Leu Leu
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                                    335
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1171
Gly Asn Ile Gly His Phe Asp Asn Glu Ile Asp Met His Ser Leu Leu
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1219
His Arg Asp Asp Val Thr Arg Thr Thr Ile Lys Pro Gln Val Asp Glu
       360
                            365
ttc acc ttc tcc acc ggt cgc tcc atc atc gtc ctg tcc gaa ggt cgc
1267
Phe Thr Phe Ser Thr Gly Arg Ser Ile Ile Val Leu Ser Glu Gly Arg
ctg ttg aac ctt ggc aac gcc acc gga cac cca tca ttt gtc atg tcc
1315
Leu Leu Asn Leu Gly Asn Ala Thr Gly His Pro Ser Phe Val Met Ser
aac tot tto goo gat cag acc att gog cag atc gaa ctg tto caa aac
Asn Ser Phe Ala Asp Gln Thr Ile Ala Gln Ile Glu Leu Phe Gln Asn
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E1.SD0010-2W0 0100843A2 LS

Asp His Ala Leu Leu Gly Asn Ile Gly His Phe Asp Asn Glu Ile Asp 355

Met His Ser Leu Leu His Arg Asp Asp Val Thr Arg Thr Thr Ile Lys 355

Pro Gln Val Asp Glu Phe Thr Phe Ser Thr Gly Asg Ser Ile Ile Val 370

Leu Ser Glu Gly Arg Leu Leu Asn Leu Gly Asg Asp Ala Thr Gly His Pro 395

Ser Phe Val Met Ser Asn Ser Phe Ala Asp Gln Thr Ile Ala Gln Ile Glu Leu Phe Gln Asn Glu Gly Gln Tyr Glu Asn Glu Val Tyr Arg Leu 425

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Met Thr Ser Asn Phe

1 5

tet tee act gte get ggt ett eet ege ate gga geg aag egt gaa etg 163 Ser Ser Thr Val Ala Gly Leu Pro Arg The Cly Ala Lye Arg Cly Lou-

15

aag ttc gcg ctc gaa ggc tac tgg aat gga tca att gaa ggt cgc gaa 211 Lys Phe Ala Leu Glu Gly Tyr Trp Asn Gly Ser Ile Glu Gly Arg Glu 25 30 35

ctt gcg cag acc gcc cgc caa ttg gtc aac act gca tcg gat tct ttg 259 Leu Ala Gln Thr Ala Arg Gln Leu Val Asn Thr Ala Ser Asp Ser Leu

tot gga ttg gat too gtt cog ttt goa gga cgt too tac tac gac gca 307 Ser Gly Leu Asp Ser Val Pro Phe Ala Gly Arg Ser Tyr Tyr Asp Ala

atg ctc gat acc gcc gct att ttg ggt gtg ctg ccg gag cgt ttt gat 355
Met Leu Asp Thr Ala Ala Ile Leu Gly Val Leu Pro Glu Arg Phe Asp
70 75 80 85

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cgt t Arg ( 150	tgc Cys	cag Gln	cag Gln	gtt Val	cgt Arg 155	ggc Gly	gtt Val	aat Asn	gcc Ala	cgc Arg 160	cct Pro	gtt Val	ctg Leu	gtt Val	ggt Gly 165	595
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tct ( Ser )	ttc Phe	gat Asp 200	act Thr	gag Glu	tgg Trp	gtt Val	cag Gln 205	atc Ile	gat Asp	gag Glu	cct Pro	gcg Ala 210	ttg Leu	gtc Val	acc Thr	739
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gtt Val	gac Asp	ttg Leu	gtc Val 265	Thr	cat His	ggc Gly	gtc Val	act Thr 270	gag Glu	ctt Leu	gct Ala	gcg Ala	tgg Trp 275	aag Lys	ggt Gly	931
gag Glu	gag Glu	ctg Leu 280	ctg Leu	gtt Val	gcg Ala	ggc Gly	atc Ile 285	Val	gat Asp	ggt Gly	cgt Arg	aac Asn 290	att Ile	tgg Trp	cgc Arg	979
acc 1027		ctg	tgt	gct	gct	ctt	gct	tcc	ctg	aag	cgc	ctg	gca	gct	cgc	
Thr	Asp 295					300					305				Arg	
1075	·								tca							
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cca atg act gtc aag tgg ttc cag tac gca cag agc ctg acc cag aag Pro Met Thr Val Lys Trp Phe Gln Tyr Ala Gln Ser Leu Thr Gln Lys 525 cat gtc aag gga atg ctc acc ggt cca gtc acc atc ctt gca tgg tcc His Val Lys Gly Met Leu Thr Gly Pro Val Thr Ile Leu Ala Trp Ser 540 ttc gtt cgc gat gat cag ccg ctg gct acc act gct gac cag gtt gca Phe Val Arg Asp Asp Gln Pro Leu Ala Thr Thr Ala Asp Gln Val Ala 555 ctg gca ctg cgc gat gaa att aac gat ctc atc gag gct ggc gcg aag 1843 Leu Ala Leu Arg Asp Glu Ile Asn Asp Leu Ile Glu Ala Gly Ala Lys 575 570 atc atc cag gtg gat gag cct gcg att cgt gaa ctg ttg ccg cta cga 1891 Ile Ile Gln Val Asp Glu Pro Ala Ile Arg Glu Leu Leu Pro Leu Arg 590 585 gac gtc gat aag cct gcc tac ctg cag tgg tcc gtg gac tcc ttc cgc 1939 Asp Val Asp Lys Pro Ala Tyr Leu Gln Trp Ser Val Asp Ser Phe Arg 600 605 ctg gcg act gcc ggc gca ccc gac gtc caa atc cac acc cac atg 1987 Leu Ala Thr Ala Gly Ala Pro Asp Asp Val Gln Ile His Thr His Met 620 615 tgc tac tcc gag ttc aac gaa gtg atc tcc tcg gtc atc gcg ttg gat 2035 Cys Tyr Ser Glu Phe Asn Glu Val Ile Ser Ser Val Ile Ala Leu Asp 640 635 630 gcc gat gtc acc acc atc gaa gca gca cgt tcc gac atg cag gtc ctc 2083 Ala Asp Val Thr Thr Ile Glu Ala Ala Arg Ser Asp Met Gln Val Leu 655 650 get get etg aaa tet tee gge tte gag ete gge gte gga eet ggt gtg 2131 Ala Ala Leu Lys Ser Ser Gly Phe Glu Leu Gly Val Gly Pro Gly Val 670 665 tgg gat atc cac tcc ccg cgc gtt cct tcc gcg cag aaa gtg gac ggt 2179 Trp Asp Ile His Ser Pro Arg Val Pro Ser Ala Gln Lys Val Asp Gly 685 680 ctc ctc gag gct gca ctg cag tcc gtg gat cct cgc cag ctg tgg gtc 2227 Leu Leu Glu Ala Ala Leu Gln Ser Val Asp Pro Arg Gln Leu Trp Val 700 705 695

BRISDOCID - 10 C10081342 LS

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tcc cta aag gtt ctc gtt gag tcc gct aag cag gct cgt gag aaa atc 2323

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Ala Ser Asp Ser Leu Ser Gly Leu Asp Ser Val Pro Phe Ala Gly Arg

Ser Tyr Tyr Asp Ala Met Leu Asp Thr Ala Ala Ile Leu Gly Val Leu 65 70 75 80

Pro Glu Arg Phe Asp Asp Ile Ala Asp His Glu Asn Asp Gly Leu Pro 85 90 95

Leu Trp Ile Asp Arg Tyr Phe Gly Ala Ala Arg Gly Thr Glu Thr Leu
100 105 110

Pro Ala Gln Ala Met Thr Lys Trp Phe Asp Thr Asn Tyr His Tyr Leu 115 120 125

Val Pro Glu Leu Ser Ala Asp Thr Arg Phe Val Leu Asp Ala Ser Ala 130 135 140

Leu Ile Glu Asp Leu Arg Cys Gln Gln Val Arg Gly Val Asn Ala Arg 145 150 155 160

Pro Val Leu Val Gly Pro Leu Thr Phe Leu Ser Leu Ala Arg Thr Thr 165 170 175

Asp Gly Ser Asn Pro Leu Asp His Leu Pro Ala Leu Phe Glu Val Tyr 180 185 190

Glu Arg Leu Ile Lys Ser Phe Asp Thr Glu Trp Val Gln Ile Asp Glu 195 200 205

Pro Ala Leu Val Thr Asp Val Ala Pro Glu Val Leu Glu Gln Val Arg 215 Ala Gly Tyr Thr Thr Leu Ala Lys Arg Asp Gly Val Phe Val Asn Thr 230 Tyr Phe Gly Ser Gly Asp Gln Ala Leu Asn Thr Leu Ala Gly Ile Gly 245 250 Leu Gly Ala Ile Gly Val Asp Leu Val Thr His Gly Val Thr Glu Leu Ala Ala Trp Lys Gly Glu Glu Leu Leu Val Ala Gly Ile Val Asp Gly Arg Asn Ile Trp Arg Thr Asp Leu Cys Ala Ala Leu Ala Ser Leu Lys Arg Leu Ala Ala Arg Gly Pro Ile Ala Val Ser Thr Ser Cys Ser Leu 310 Leu His Val Pro Tyr Thr Leu Glu Ala Glu Asn Ile Glu Pro Glu Val Arg Asp Trp Leu Ala Phe Gly Ser Glu Lys Ile Thr Glu Val Lys Leu 345 Leu Ala Asp Ala Leu Ala Gly Asn Ile Asp Ala Ala Ala Phe Asp Ala Ala Ser Ala Ala Ile Ala Ser Arg Arg Thr Ser Pro Arg Thr Ala Pro Ile Thr Gln Glu Leu Pro Gly Arg Ser Arg Gly Ser Phe Asp Thr Arg Val Thr Leu Gln Glu Lys Ser Leu Glu Leu Pro Ala Leu Pro Thr Thr Thr Ile Gly Ser Phe Pro Gln Thr Pro Ser Ile Arg Ser Ala Arg Ala Arg Leu Arg Lys Glu Ser Ile Thr Leu Glu Gln Tyr Glu Glu Ala Met Arg Glu Glu Ile Asp Leu Val Ile Ala Lys Gln Glu Glu Leu Gly Leu Asp Val Leu Val His Gly Glu Pro Glu Arg Asn Asp Met Val Gln Tyr Phe Ser Glu Leu Leu Asp Gly Phe Leu Ser Thr Ala Asn Gly Trp Val Gln Ser Tyr Gly Ser Arg Cys Val Arg Pro Pro Val Leu Phe Gly Asn 505 Val Ser Arg Pro Ala Pro Met Thr Val Lys Trp Phe Gln Tyr Ala Gln 515 520 Ser Leu Thr Gln Lys His Val Lys Gly Met Leu Thr Gly Pro Val Thr

ENSTROCK AWO 0100843A2 1 >

535 530 Ile Leu Ala Trp Ser Phe Val Arg Asp Asp Gln Pro Leu Ala Thr Thr 550 555 Ala Asp Gln Val Ala Leu Ala Leu Arg Asp Glu Ile Asn Asp Leu Ile 565 570 Glu Ala Gly Ala Lys Ile Ile Gln Val Asp Glu Pro Ala Ile Arg Glu 585 Leu Leu Pro Leu Arg Asp Val Asp Lys Pro Ala Tyr Leu Gln Trp Ser 600 Val Asp Ser Phe Arg Leu Ala Thr Ala Gly Ala Pro Asp Asp Val Gln 615 Ile His Thr His Met Cys Tyr Ser Glu Phe Asn Glu Val Ile Ser Ser Val Ile Ala Leu Asp Ala Asp Val Thr Thr Ile Glu Ala Ala Arg Ser 645 650 Asp Met Gln Val Leu Ala Ala Leu Lys Ser Ser Gly Phe Glu Leu Gly Val Gly Pro Gly Val Trp Asp Ile His Ser Pro Arg Val Pro Ser Ala 680 Gln Lys Val Asp Gly Leu Leu Glu Ala Ala Leu Gln Ser Val Asp Pro Arg Gln Leu Trp Val Asn Pro Asp Cys Gly Leu Lys Thr Arg Gly Trp 710 Pro Glu Val Glu Ala Ser Leu Lys Val Leu Val Glu Ser Ala Lys Gln Ala Arg Glu Lys Ile Gly Ala Thr Ile

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	gcg Ala															259
	gga Gly 55															307
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	atc Ile															403
	ttt Phe															451
	aag Lys															499
	gat Asp 135															547
	tgc Cys															595
	ctg Leu															643
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ttg Leu 230	gct Ala	aag Lys	cgc Arg	gat Asp	ggc Gly 235	gtg Val	ttt Phe	gtc Val	aat Asn	act Thr 240	tac Tyr	ttc Phe	ggc Gly	tct Ser	ggc Gly 245	835
gat Asp	cag Gln	gcg Ala	ctg Leu	aac Asn 250	act Thr	ctt Leu	gcg Ala	ggc Gly	atc Ile 255	ggc Gly	ctt Leu	Gly	gcg Ala	att Ile 260	ggc Gly	883

gtt gac Val Asp															931
gag gag Glu Glu															9 <b>79</b>
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Thr Asp 295	Leu	Cys	Ala	Ala	Leu 300	Ala	Ser	Leu	Lys	Arg 305	Leu	Ala	Ala	Arg	
ggc cca 1075	atc	gca	gtg	tct	acc	tct	tgt	tca	ctg	ctg	cac	gtt	cct	tac	
Gly Pro	Ile	Ala	Val	Ser 315	Thr	Ser	Cys	Ser	Leu 320	Leu	His	Val	Pro	Tyr 325	
acc ctc 1123	gag	gct	gag	aac	att	gag	cct	gag	gtc	cgc	gac	tgg	ctt	gcc	
Thr Leu	Glu	Ala	Glu 330	Asn	Ile	Glu	Pro	Glu 335	Val	Arg	Asp	Trp	Leu 340	Ala	
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Phe Gly	Ser	Glu 345	Lys	Ile	Thr	Glu	Val 350	Lys	Leu	Leu	Ala	Asp 355	Ala	Leu	
gcc ggc 1219	aac	atc	gac	gcg	gct	gcg	ttc	gat	gcg	gcg	tcc	gca	gca	att	
Ala Gly	Asn 360	Ile	Asp	Ala	Ala	Ala 365	Phe	Asp	Ala	Ala	Ser 370	Ala	Ala	Ile	
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Ala Ser 375	Arg	Arg	Thr	Ser	Pro 380	Arg	Thr	Ala	Pro	Ile 385	Thr	Gln	Glu	Leu	
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PRISONOLO SALO CARGERIZAS I S

Ile Glu Gly Arg Glu Leu Ala Gln Thr Ala Arg Gln Leu Val Asn Thr Ala Ser Asp Ser Leu Ser Gly Leu Asp Ser Val Pro Phe Ala Gly Arg Ser Tyr Tyr Asp Ala Met Leu Asp Thr Ala Ala Ile Leu Gly Val Leu Pro Glu Arg Phe Asp Asp Ile Ala Asp His Glu Asn Asp Gly Leu Pro Leu Trp Ile Asp Arg Tyr Phe Gly Ala Ala Arg Gly Thr Glu Thr Leu 105 Pro Ala Gln Ala Met Thr Lys Trp Phe Asp Thr Asn Tyr His Tyr Leu Val Pro Glu Leu Ser Ala Asp Thr Arg Phe Val Leu Asp Ala Ser Ala 135 Leu Ile Glu Asp Leu Arg Cys Gln Gln Val Arg Gly Val Asn Ala Arg Pro Val Leu Val Gly Pro Leu Thr Phe Leu Ser Leu Ala Arg Thr Thr 170 Asp Gly Ser Asn Pro Leu Asp His Leu Pro Ala Leu Phe Glu Val Tyr Glu Arg Leu Ile Lys Ser Phe Asp Thr Glu Trp Val Gln Ile Asp Glu 200 Pro Ala Leu Val Thr Asp Val Ala Pro Glu Val Leu Glu Gln Val Arg Ala Gly Tyr Thr Thr Leu Ala Lys Arg Asp Gly Val Phe Val Asn Thr 230 235 Tyr Phe Gly Ser Gly Asp Gln Ala Leu Asn Thr Leu Ala Gly Ile Gly Leu Gly Ala Ile Gly Val Asp Leu Val Thr His Gly Val Thr Ala Ala Trp Lys Gly Glu Glu Leu Leu Val Ala Gly Ile Val Asp Gly Arg Asn Ile Trp Arg Thr Asp Leu Cys Ala Ala Leu Ala Ser Leu Lys 295 Arg Leu Ala Ala Arg Gly Pro Ile Ala Val Ser Thr Ser Cys Ser Leu 315 Leu His Val Pro Tyr Thr Leu Glu Ala Glu Asn Ile Glu Pro Glu Val 330 Arg Asp Trp Leu Ala Phe Gly Ser Glu Lys Ile Thr Glu Val Lys Leu Leu Ala Asp Ala Leu Ala Gly Asn Ile Asp Ala Ala Ala Phe Asp Ala

360 365 355 Ala Ser Ala Ala Ile Ala Ser Arg Arg Thr Ser Pro Arg Thr Ala Pro 380 375 Ile Thr Gln Glu Leu Pro Gly Arg Ser Arg Gly Ser Phe Asp Thr Arg Val Thr Leu Gln Glu Lys Ser Leu Glu Leu Pro Ala Leu Pro Thr Thr 410 Thr Ile Gly Ser Phe Pro Gln Thr Pro Ser Ile Arg Ser Ala Arg Ala 425 420 Arg Leu Arg Lys Glu Ser Ile Thr Leu Glu Gln Tyr Glu Glu Ala Met 440 Arg Glu Glu Ile Asp Leu Val Ile Ala Lys Gln Glu Glu Leu Gly Leu 455 450 Asp Val Leu Val His Gly Glu Pro Glu Arg Asn Asp Met Val Gln Tyr 475 470 Phe Ser Glu Leu Leu Asp Gly Phe Leu Ser Thr Ala Asn Gly Trp Val 490 485 Gln Ser Tyr Gly Ser Arg Cys Val Arg Pro Pro Val Leu Phe Gly Asn 505 500 Val Ser Arg Pro Ala Pro Met Thr Val Lys Trp Phe Gln Tyr Ala Gln 520 515 Ser Leu Thr Gln Lys His Val Lys Gly Met Leu Thr Gly Pro Val Thr 535 Ile Leu Ala Trp Ser Phe Val Arg Asp Asp Gln Pro Leu Ala Thr Thr 550 545 Ala Asp Gln Val Ala Leu Ala Leu Arg Asp Glu Ile Asn Asp Leu Ile 570 565 Glu Ala Gly Ala Lys Ile Ile Gln Val Asp Glu Pro Ala Ile Arg Glu 580 585 Leu Leu Pro Ala Thr Arg Arg Arg 595 <210> 229 <211> 603 <212> DNA <213> Corynebacterium glutamicum <220> <221> CDS <222> (101)..(580) <223> FRXA02086 <400> 229

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tgg tcc gtg gac tcc ttc cgc ctg gcg act gcc ggc gca ccc gac gac 21 Trp Ser Val Asp Ser Phe Arg Leu Ala Thr Ala Gly Ala Pro Asp Asp 25 30 35	11
gtc caa atc cac acc cac atg tgc tac tcc gag ttc aac gaa gtg atc 25 Val Gln Ile His Thr His Met Cys Tyr Ser Glu Phe Asn Glu Val Ile 40 45 50	59
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Phe Asn Glu Val Ile Ser Ser Val Ile Ala Leu Asp Ala Asp Val Thr 55 Val Ile Ala Leu Asp Ala Asp Val Thr 66 Val Glu Ala Ala Arg 70 Val Glu Pro 75 Val Leu Ala Ala Leu Lys 80 Ser Ser Gly Phe Glu Leu Gly Val Gly Pro 90 Gly Val Trp Asp Ile His 95 Val Ala 100 Val 105 Val Asp Gly Leu Leu Glu Ala 110 Glu Ala 110 Gly Ileu Leu Gli Ser Val Asp Pro Arg 120 Gln Leu Trp Val Asp Pro Asp Cys 130 Val Ileu Lys Thr Arg Gly Trp Pro Glu Val Glu Ala Ser Leu Lys Val 145 Val Glu Ser Ala Lys Gln Ala Arg Glu Lys Ile Gly Ala Thr Ile 160

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-	_				cct Pro											451
					ttg Leu											499
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					ggc Gly 155											595
					aac Asn											643
_					tct Ser	_	_	_	_							691
					gtc Val											739
					gat Asp											787
					tgg Trp 235											835
															αca	883
Asp	Tyr	Leu	Asp	Trp 250	Ile	Gly	Thr	Arg	11e 255	Asp	Ala	Ile	Asn	Ser 260	Ala	
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					cac His											979
		atc	ctg	cgc	gca	gag	gtc	ggt	ggc	ttc	tcc	ttc	gaa	ggc	gca	
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Pro Glu Gly Ser Val Ile Tyr Pro Gly Val Val Ser His Ser Ile Asn 330 335 340

gct gtg gag cac cca cgc ctg gtt gct gat cgt atc gtt cag ttc gcc 1171

Ala Val Glu His Pro Arg Leu Val Ala Asp Arg Ile Val Gln Phe Ala 345 350 355

aag ctt g<br/>tt ggc cct gag aac gtc att gcg tcc act gac tg<br/>t ggt ctg  $1219\,$ 

ggc gga cgt ctg cat tcc cag atc gca tgg gca aag ctg gag tcc cta 1267

Gly Gly Arg Leu His Ser Gln Ile Ala Trp Ala Lys Leu Glu Ser Leu 375 380 385

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<400> 232

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Thr Pro Glu Leu Leu Asp Ala Asn Ile Lys Arg Ser Asn Gly Glu Ile 20 25 30

Gly Glu Glu Phe Phe Gln Ile Leu Gln Ser Ser Val Asp Asp Val

Ile Lys Arg Gln Val Asp Leu Gly Ile Asp Ile Leu Asn Glu Gly Glu 50 55 60

Tyr Gly His Val Thr Ser Gly Ala Val Asp Phe Gly Ala Trp Trp Asn 65 70 75 80

Tyr Ser Phe Thr Arg Leu Gly Gly Leu Thr Met Thr Asp Thr Asp Arg  $85 \hspace{1cm} 90 \hspace{1cm} 95$ 

Trp Ala Ser Gln Glu Ala Val Arg Ser Thr Pro Gly Asn Ile Glu Leu 100 105 110

Thr Ser Phe Ser Asp Arg Arg Asp Arg Ala Leu Phe Ser Glu Ala Tyr
115 120 125

Glu Asp Pro Val Ser Gly Ile Phe Thr Gly Arg Ala Ser Val Gly Asn Pro Glu Phe Thr Gly Pro Ile Thr Tyr Ile Gly Gln Glu Glu Thr Gln 150 Thr Asp Val Asp Leu Leu Lys Lys Gly Met Asn Ala Ala Gly Ala Thr 170 Asp Gly Phe Val Ala Ala Leu Ser Pro Gly Ser Ala Ala Arg Leu Thr 185 Asn Lys Phe Tyr Asp Thr Asp Glu Glu Val Val Ala Ala Cys Ala Asp Ala Leu Ser Gln Glu Tyr Lys Ile Ile Thr Asp Ala Gly Leu Thr Val 215 Gln Leu Asp Ala Pro Asp Leu Ala Glu Ala Trp Asp Gln Ile Asn Pro 230 Glu Pro Ser Val Lys Asp Tyr Leu Asp Trp Ile Gly Thr Arg Ile Asp 250 Ala Ile Asn Ser Ala Val Lys Gly Leu Pro Lys Glu Gln Thr Arg Leu His Ile Cys Trp Gly Ser Trp His Gly Pro His Val Thr Asp Ile Pro 280 Phe Gly Asp Ile Ile Gly Glu Ile Leu Arg Ala Glu Val Gly Phe Ser Phe Glu Gly Ala Ser Pro Arg His Ala His Glu Trp Arg Val Trp 310 315 Glu Glu Asn Lys Leu Pro Glu Gly Ser Val Ile Tyr Pro Gly Val Val 325 330 Ser His Ser Ile Asn Ala Val Glu His Pro Arg Leu Val Ala Asp Arg 345 Ile Val Gln Phe Ala Lys Leu Val Gly Pro Glu Asn Val Ile Ala Ser Thr Asp Cys Gly Leu Gly Gly Arg Leu His Ser Gln Ile Ala Trp Ala 375 380 Lys Leu Glu Ser Leu Val Glu Gly Ala Arg Ile Ala Ser Lys Glu Leu 390 395 Phe

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45

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345

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